

## Protein Sequence Searches - February 2005

All of the sequence databases on ABSS have recently been updated.

- Please note that the curators of the UniProt database have purged some temporary accession numbers from the most recent version of UniProt. These sequences have been assigned new permanent accession numbers. The new UniProt record may not contain the previous temporary accession number.
- If you encounter an accession number from an older search run against UniProt (results file extension **.rup**) that can no longer be found in the database, the permanent record with the new accession number can be found by searching the old accession number in the UniProt Protein Archive database (UniPARC) at:

<http://www.pir.uniprot.org/database/archive.shtml>

If you have any questions regarding this information or your results, please contact any STIC searcher.

**When submitting sequence search results for scanning into IFW, please include a copy of this attachment to assist any future Examiners or members of the public who may encounter UniProt temporary accession numbers.**

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160352

From: Swope, Sheridan  
Sent: Monday, July 25, 2005 2:40 PM  
To: STIC-Biotech/ChemLib  
Subject: 10/726,967

For 10/726,967, pls search:

(A)--Claim 1(18)

SID 3: oligo search against the NT & AA data bases.

(B)

SID 52 against the NT & AA data bases.

For any hits that are 100% identical, pls align with:  
residues 22-41 of SID 1,

CRF

If you have questions, pls ask me or David Schreiber.

Thanks

Sheridan Swope, Ph.D.  
Patent Examiner, AU 1656  
Recombinant Enzymes  
571-272-0943 (voice)  
E02B71 Remsen Bld (Office)  
E03C70 Remsen Bld (Mailbox)

\*\*\*\*\*  
STAFF USE ONLY

Searcher: \_\_\_\_\_  
Searcher Phone: 2- \_\_\_\_\_  
Date Searcher Picked up: \_\_\_\_\_  
Date Completed: \_\_\_\_\_  
Searcher Prep/Rev. Time: \_\_\_\_\_  
Online Time: \_\_\_\_\_

\*\*\*\*\*  
Type of Search

NA#: \_\_\_\_\_ AA#: \_\_\_\_\_  
Interference: \_\_\_\_\_ SPDI: \_\_\_\_\_  
S/L: \_\_\_\_\_ Oligomer: \_\_\_\_\_  
Encode/Transl: \_\_\_\_\_  
Structure#: \_\_\_\_\_ Text: \_\_\_\_\_  
Inventor: \_\_\_\_\_ Litigation: \_\_\_\_\_

\*\*\*\*\*  
Vendors and cost where applicable

STN: \_\_\_\_\_  
DIALOG: \_\_\_\_\_  
QUESTEL/ORBIT: \_\_\_\_\_  
LEXIS/NEXIS: \_\_\_\_\_  
SEQUENCE SYSTEM: \_\_\_\_\_  
WWW/Internet: \_\_\_\_\_  
Other(Specify): \_\_\_\_\_

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(uspto)



GenCore version 5.1.6  
Copyright (c) 1993 - 2005 CompuGen Ltd.

OM protein - protein search, using sw model

Run on: July 26, 2005, 16:28:49 ; Search time 43 Seconds  
(without alignments)  
27.776 Million cell updates/sec

Title: US-10-726-967A-3

Perfect score: 16

Sequence: 1 TQHGRLPLRSGLGCA 16

Scoring table: OLIGO

Gapop 60.0 , Gapext 60.0

Searched: 513545 seqs, 74649064 residues

Word size : 0

Total number of hits satisfying chosen parameters: 513545

Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Listing first 45 summaries

Database : Issued Patents AA:\*

1: /cgn2\_6/prodata/1/1aa/5A\_COMB.pep:\*\n2: /cgn2\_6/prodata/1/1aa/5B\_COMB.pep:\*\n3: /cgn2\_6/prodata/1/1aa/6A\_COMB.pep:\*\n4: /cgn2\_6/prodata/1/1aa/6B\_COMB.pep:\*\n5: /cgn2\_6/prodata/1/1aa/6C\_COMB.pep:\*\n6: /cgn2\_6/prodata/1/1aa/6ackfile1.pep:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	16	100.0	24	4	US-09-724-566A-47
2	16	100.0	419	4	US-09-724-566A-57
3	16	100.0	419	4	US-09-471-669A-57
4	16	100.0	420	4	US-09-724-566A-60
5	16	100.0	420	4	US-09-471-669A-60
6	16	100.0	428	4	US-09-548-372D-51
7	16	100.0	428	4	US-09-548-367D-51
8	16	100.0	428	4	US-09-551-853D-51
9	16	100.0	428	4	US-09-416-901B-51
10	16	100.0	428	4	US-09-548-376D-51
11	16	100.0	428	4	US-09-794-927A-51
12	16	100.0	428	4	US-09-548-373D-51
13	16	100.0	428	4	US-09-795-847B-51
14	16	100.0	428	4	US-09-869-414-51
15	16	100.0	428	4	US-09-548-366F-51
16	16	100.0	428	4	US-09-548-368D-51
17	16	100.0	428	4	US-09-794-925A-51
18	16	100.0	431	4	US-09-724-566A-74
19	16	100.0	431	4	US-09-471-669A-74
20	16	100.0	433	4	US-09-548-372D-26
21	16	100.0	433	4	US-09-548-367D-26
22	16	100.0	433	4	US-09-551-853D-26
23	16	100.0	433	4	US-09-416-901B-26
24	16	100.0	433	4	US-09-548-376D-26
25	16	100.0	433	4	US-09-794-927A-26
26	16	100.0	433	4	US-09-548-373D-26
27	16	100.0	433	4	US-09-795-847B-26

28	16	100.0	433	4	US-09-869-414-26	Sequence 26, Appl
29	16	100.0	433	4	US-09-548-366F-26	Sequence 26, Appl
30	16	100.0	433	4	US-09-548-368D-26	Sequence 26, Appl
31	16	100.0	433	4	US-09-794-925A-26	Sequence 26, Appl
32	16	100.0	433	4	US-09-806-194A-26	Sequence 26, Appl
33	16	100.0	434	4	US-09-548-372D-53	Sequence 53, Appl
34	16	100.0	434	4	US-09-548-367D-53	Sequence 53, Appl
35	16	100.0	434	4	US-09-551-853D-53	Sequence 53, Appl
36	16	100.0	434	4	US-09-416-901B-53	Sequence 53, Appl
37	16	100.0	434	4	US-09-548-376D-53	Sequence 53, Appl
38	16	100.0	434	4	US-09-794-927A-53	Sequence 53, Appl
39	16	100.0	434	4	US-09-548-372D-53	Sequence 53, Appl
40	16	100.0	434	4	US-09-795-847B-53	Sequence 53, Appl
41	16	100.0	434	4	US-09-869-414-53	Sequence 53, Appl
42	16	100.0	434	4	US-09-548-366F-53	Sequence 53, Appl
43	16	100.0	434	4	US-09-548-368D-53	Sequence 53, Appl
44	16	100.0	434	4	US-09-794-925A-53	Sequence 53, Appl
45	16	100.0	446	4	US-09-548-372D-22	Sequence 22, Appl

## ALIGNMENTS

```
RESULT 1
US-09-724-566A-47
; Sequence 47, Application US/09724566A
; Patent No. 6627739
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basl, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6627739mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; TITLE OF INVENTION: Beta-Secretase Enzyme Compositions and
; FILE REFERENCE: 228-US-NEWC2
; CURRENT FILING DATE: 2000-11-28
; PRIOR APPLICATION NUMBER: US 09/724,566A
; PRIOR FILING DATE: 2000-02-10
; PRIOR APPLICATION NUMBER: 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 104
; SOFTWARE: FastSeq for Windows Version 4.0
; LENGTH: 24
; SEQ ID NO 47
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-724-566A-47

Query Match      100.0%; Score 16; DB 4; Length 24;
Best Local Similarity 100.0%; Pred. No. 8e-10;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Cy      1 TQHGRLPLRSGLGCA 16
Db      1 TQHGRLPLRSGLGCA 16

RESULT 2
US-09-724-566A-57
; Sequence 57, Application US/09724566A
; Patent No. 6627739
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
```

```

; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6627739mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuono, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; TITLE OF INVENTION: Beta-Secretase Enzyme Compositions and
; FILE REFERENCE: 228-US-NEWC2
; CURRENT APPLICATION NUMBER: US/09/724,566A
; PRIOR FILING DATE: 2000-11-28
; PRIOR APPLICATION NUMBER: US 09/501,708
; PRIOR FILING DATE: 2000-02-10
; PRIOR APPLICATION NUMBER: 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 104
; SOFTWARE: FaetsEQ for Windows Version 4.0
; SEQ ID NO 57
; LENGTH: 419
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-724-566A-57
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Query Match      100.0%; Score 16; DB 4; Length 419;
Best Local Similarity 100.0%; Pred. No. 1,1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
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```

QY      1  TONGIRLPLRSGLGGA 16
Db      22 TONGIRLPLRSGLGGA 37
```

```

RESULT 3
US-09-471-669A-57
; Sequence 57, Application US/09471669A
; Patent No. 6830918
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6830918mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuono, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS
; FILE REFERENCE: 015270-006430US
; CURRENT APPLICATION NUMBER: US/09/471,669A
; PRIOR FILING DATE: 1999-12-24
; PRIOR APPLICATION NUMBER: US 60/114,408
; PRIOR FILING DATE: 1998-12-31
; PRIOR APPLICATION NUMBER: US 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: US 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 108
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 57
; LENGTH: 419
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-471-669A-57
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Query Match      100.0%; Score 16; DB 4; Length 419;
Best Local Similarity 100.0%; Pred. No. 1,1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
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```

QY      1  TONGIRLPLRSGLGGA 16
Db      22 TONGIRLPLRSGLGGA 37
```

```

RESULT 4
US-09-724-566A-60
; Sequence 60, Application US/09724566A
; Patent No. 6627739
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6627739mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuono, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; TITLE OF INVENTION: Beta-Secretase Enzyme Compositions and
; FILE REFERENCE: 228-US-NEWC2
; CURRENT APPLICATION NUMBER: US/09/724,566A
; PRIOR FILING DATE: 2000-11-28
; PRIOR APPLICATION NUMBER: US 09/501,708
; PRIOR FILING DATE: 2000-02-10
; PRIOR APPLICATION NUMBER: 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 104
; SOFTWARE: FaetsEQ for Windows Version 4.0
; SEQ ID NO 60
; LENGTH: 420
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-724-566A-60
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Query Match      100.0%; Score 16; DB 4; Length 420;
Best Local Similarity 100.0%; Pred. No. 1,1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
```

```

QY      1  TONGIRLPLRSGLGGA 16
Db      22 TONGIRLPLRSGLGGA 37
```

```

RESULT 5
US-09-471-669A-60
; Sequence 60, Application US/09471669A
; Patent No. 6830918
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6830918mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuono, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS
; FILE REFERENCE: 015270-006430US
; CURRENT APPLICATION NUMBER: US/09/471,669A
```

```

; CURRENT FILING DATE: 1999-12-24
; PRIOR APPLICATION NUMBER: US 60/114,408
; PRIOR FILING DATE: 1998-12-31
; PRIOR APPLICATION NUMBER: US 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: US 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 108
; SOFTWARE: Patentn Ver. 2.1
; SEQ ID NO 60
; LENGTH: 420
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-471-669A-60

Query Match          100.0%; Score 16; DB 4; Length 420;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 TOHGIRLPLRSGGCA 16
        |||||||
        22 TOHGIRLPLRSGGCA 37

Db

RESULT 6
US-09-548-372D-51
; Sequence 51, Application US/09548372D
; Patent No. 6420534
; GENERAL INFORMATION:
; APPLICANT: GURNEY ET AL.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES
; FILE REFERENCE: 29915/62801
; CURRENT FILING DATE: 2000-04-12
; PRIOR APPLICATION NUMBER: US 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentn version 3.1
; SEQ ID NO 51
; LENGTH: 428
; TYPE: PRT
; ORGANISM: Artificial sequence
; FEATURE:
; OTHER INFORMATION: Hu-Asp2 (b) delta TM
US-09-548-372D-51

Query Match          100.0%; Score 16; DB 4; Length 428;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 TOHGIRLPLRSGGCA 16
        |||||||
        22 TOHGIRLPLRSGGCA 37

Db

RESULT 7
US-09-548-367D-51
; Sequence 51, Application US/09548367D
; Patent No. 6440698
; GENERAL INFORMATION:
; APPLICANT: GURNEY ET AL.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES
; FILE REFERENCE: 29915/6280H
; CURRENT FILING DATE: 2000-04-12
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```

; PRIOR APPLICATION NUMBER: US 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentn version 3.1
; SEQ ID NO 51
; LENGTH: 428
; TYPE: PRT
; ORGANISM: Artificial sequence
; FEATURE:
; OTHER INFORMATION: Hu-Asp2 (b) delta TM
US-09-548-367D-51

Query Match          100.0%; Score 16; DB 4; Length 428;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 TOHGIRLPLRSGGCA 16
        |||||||
        22 TOHGIRLPLRSGGCA 37

Db

RESULT 8
US-09-551-853D-51
; Sequence 51, Application US/09551853D
; Patent No. 6500667
; GENERAL INFORMATION:
; APPLICANT: GURNEY ET AL.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES
; FILE REFERENCE: 29915/6280L
; CURRENT FILING DATE: 2000-04-18
; PRIOR APPLICATION NUMBER: US 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentn version 3.1
; SEQ ID NO 51
; LENGTH: 428
; TYPE: PRT
; ORGANISM: Artificial sequence
; FEATURE:
; OTHER INFORMATION: Hu-Asp2 (b) delta TM
US-09-551-853D-51

Query Match          100.0%; Score 16; DB 4; Length 428;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 TOHGIRLPLRSGGCA 16
        |||||||
        22 TOHGIRLPLRSGGCA 37

Db

RESULT 9
US-09-416-901B-51
; Sequence 51, Application US/09416901B
; Patent No. 669671
; GENERAL INFORMATION:
; APPLICANT: GURNEY ET AL.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES
```

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FILE REFERENCE: 29915/6280A
CURRENT APPLICATION NUMBER: US/09/416,901B
CURRENT FILING DATE: 1999-10-13
PRIORITY APPLICATION NUMBER: US 60/155,493
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: US 09/404,133
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: PCT/US99/20881
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: US 60/101,594
PRIORITY FILING DATE: 1998-09-24
NUMBER OF SEQ ID NOS: 72
SOFTWARE: Patentin version 3.1
SEQ ID NO 51
LENGTH: 428
TYPE: PRT
ORGANISM: Artificial sequence
FEATURE:
OTHER INFORMATION: Hu-Asp2(b) delta TM
US-09-416-901B-51
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```
Query Match          100.0%; Score 16; DB 4; Length 428;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
```

```
Qy      1 TOHGIRLPLRSGGGA 16
        |||
Db      22 TOHGIRLPLRSGGGA 37
```

```
RESULT 10
US-09-548-376D-51
Sequence 51, Application US/09548376D
Patent No. 6706485
GENERAL INFORMATION:
APPLICANT: GURNEY ET AL.
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR
TITLE OF INVENTION: AND USES
FILE REFERENCE: 29915/6280F
CURRENT APPLICATION NUMBER: US/09/548,376D
CURRENT FILING DATE: 2000-04-12
PRIORITY APPLICATION NUMBER: US 60/155,493
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: US 09/404,133
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: PCT/US99/20881
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: US 60/101,594
PRIORITY FILING DATE: 1998-09-24
NUMBER OF SEQ ID NOS: 73
SOFTWARE: Patentin version 3.1
SEQ ID NO 51
LENGTH: 428
TYPE: PRT
ORGANISM: Artificial sequence
FEATURE:
OTHER INFORMATION: Hu-Asp2(b) delta TM
US-09-548-376D-51
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```
Query Match          100.0%; Score 16; DB 4; Length 428;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
```

```
Qy      1 TOHGIRLPLRSGGGA 16
        |||
Db      22 TOHGIRLPLRSGGGA 37
```

```
RESULT 11
US-09-794-927A-51
Sequence 51, Application US/09794927A
Patent No. 6727074
```

```
GENERAL INFORMATION:
APPLICANT: Gurney et al.
TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses
TITLE OF INVENTION: THEREFOR
FILE REFERENCE: 29915/6280FG
CURRENT APPLICATION NUMBER: US/09/794,927A
CURRENT FILING DATE: 2001-02-27
PRIORITY APPLICATION NUMBER: 09/416,901
PRIORITY FILING DATE: 1999-10-13
PRIORITY APPLICATION NUMBER: 60/155,493
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: 09/404,133
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: PCT/US99/20881
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: 60/101,594
PRIORITY FILING DATE: 1998-09-24
NUMBER OF SEQ ID NOS: 74
SOFTWARE: Patentin Ver. 2.0
SEQ ID NO 51
LENGTH: 428
TYPE: PRT
ORGANISM: Artificial sequence
FEATURE:
OTHER INFORMATION: Hu-Asp2(b) delta TM
US-09-794-927A-51
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Query Match          100.0%; Score 16; DB 4; Length 428;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
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```
Qy      1 TOHGIRLPLRSGGGA 16
        |||
Db      22 TOHGIRLPLRSGGGA 37
```

```
RESULT 12
US-09-548-373D-51
Sequence 51, Application US/09548373D
Patent No. 6737510
GENERAL INFORMATION:
APPLICANT: GURNEY ET AL.
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES
TITLE OF INVENTION: THEREOF
FILE REFERENCE: 29915/6280B
CURRENT APPLICATION NUMBER: US/09/548,373D
CURRENT FILING DATE: 2000-04-12
PRIORITY APPLICATION NUMBER: US 60/155,493
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: US 09/404,133
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: PCT/US99/20881
PRIORITY FILING DATE: 1999-09-23
PRIORITY APPLICATION NUMBER: US 60/101,594
PRIORITY FILING DATE: 1998-09-24
NUMBER OF SEQ ID NOS: 73
SOFTWARE: Patentin version 3.1
SEQ ID NO 51
LENGTH: 428
TYPE: PRT
ORGANISM: Artificial sequence
FEATURE:
OTHER INFORMATION: Hu-Asp2(b) delta TM
US-09-548-373D-51
```

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Query Match          100.0%; Score 16; DB 4; Length 428;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
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```
Qy      1 TOHGIRLPLRSGGGA 16
        |||
Db      22 TOHGIRLPLRSGGGA 37
```

```
RESULT 13
US-09-795-847B-51
; Sequence 51, Application US/09795847B
; Patent No. 6753163
; GENERAL INFORMATION:
; APPLICANT: Guiney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrichson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; FILE REFERENCE: 28341/6280DE
; CURRENT APPLICATION NUMBER: US/09/795,847B
; CURRENT FILING DATE: 2001-02-28
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 74
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 51
; LENGTH: 428
; TYPE: PRT
; ORGANISM: Artificial sequence
; FEATURE:
; OTHER INFORMATION: Hu-Asp2 (b) delta TM
US-09-795-847B-51

Query Match          100.0%; Score 16; DB 4; Length 428;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 TOHGIRLPLRSGIGGA 16
        |||||||
Db      22 TOHGIRLPLRSGIGGA 37

RESULT 14
US-09-869-414-51
; Sequence 51, Application US/09869414
; Patent No. 6790610
; GENERAL INFORMATION:
; APPLICANT: Bienkowski et al.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; FILE REFERENCE: 28341/6280M
; CURRENT APPLICATION NUMBER: US/09/869,414
; CURRENT FILING DATE: 2001-06-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 51
; LENGTH: 428
; TYPE: PRT
; ORGANISM: Artificial Sequence
; FEATURE:
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```
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
; OTHER INFORMATION: delta TM
US-09-869-414-51

Query Match          100.0%; Score 16; DB 4; Length 428;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 TOHGIRLPLRSGIGGA 16
        |||||||
Db      22 TOHGIRLPLRSGIGGA 37

RESULT 15
US-09-548-366F-51
; Sequence 51, Application US/09548366F
; Patent No. 6797487
; GENERAL INFORMATION:
; APPLICANT: GURNEY ET AL.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES
; FILE REFERENCE: 29915/6280J
; CURRENT APPLICATION NUMBER: US/09/548,366F
; CURRENT FILING DATE: 2000-04-12
; PRIOR APPLICATION NUMBER: US 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentin version 3.1
; SEQ ID NO 51
; LENGTH: 428
; TYPE: PRT
; ORGANISM: Artificial sequence
; FEATURE:
; OTHER INFORMATION: Hu-Asp2 (b) delta TM
US-09-548-366F-51

Query Match          100.0%; Score 16; DB 4; Length 428;
Best Local Similarity 100.0%; Pred. No. 1.1e-08;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 TOHGIRLPLRSGIGGA 16
        |||||||
Db      22 TOHGIRLPLRSGIGGA 37

Search completed: July 26, 2005, 16:38:50
Job time : 44 secs
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GenCore version 5.1.6  
Copyright (c) 1993 - 2005 CompuGen Ltd.

OM protein - protein search, using sw model

Run on: July 26, 2005, 16:24:19 ; Search time 166 Seconds  
(without alignments)  
37.278 Million cell updates/sec

Title: US-10-726-967A-3

Perfect score: 16

Sequence: 1 TQHGIRLPKRSGLGGA 16

Scoring table: OLIGO

Gapop 60.0 , Gapext 60.0

Searched: 2105692 seqs, 386760381 residues

Word size : 0

Total number of hits satisfying chosen parameters: 2105692

Minimum DB seq length: 0

Maximum DB seq length: 200000000

Post-processing: listing first 45 summaries

Database : A\_Geneseq\_16Dec04:\*

1: geneeqp1980a:\*  
2: geneeqp1990a:\*  
3: geneeqp2000a:\*  
4: geneeqp2001a:\*  
5: geneeqp2002a:\*  
6: geneeqp2003a:\*  
7: geneeqp2003b:\*  
8: geneeqp2004a:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	16	100.0	16	8 ADP83877	Adp83877 Human BAC
2	16	100.0	425	7 ADC81580	Adc81580 Beta-secr
3	16	100.0	428	4 AAU07219	AAu07219 Human asp
4	16	100.0	428	4 AAU07219	AAu07219 Human asp
5	16	100.0	428	4 AAU07219	AAu07219 Human asp
6	16	100.0	428	4 AAU07219	AAu07219 Human asp
7	16	100.0	428	4 AAU07219	AAu07219 Human asp
8	16	100.0	428	4 AAU07219	AAu07219 Human asp
9	16	100.0	428	4 AAU07219	AAu07219 Human asp
10	16	100.0	428	4 AAU07219	AAu07219 Human asp
11	16	100.0	428	4 AAU07219	AAu07219 Human asp
12	16	100.0	428	4 AAU07219	AAu07219 Human asp
13	16	100.0	428	4 AAU07219	AAu07219 Human asp
14	16	100.0	428	4 AAU07219	AAu07219 Human asp
15	16	100.0	428	4 AAU07219	AAu07219 Human asp
16	16	100.0	428	4 AAU07219	AAu07219 Human asp
17	16	100.0	428	4 AAU07219	AAu07219 Human asp
18	16	100.0	428	4 AAU07219	AAu07219 Human asp
19	16	100.0	428	4 AAU07219	AAu07219 Human asp
20	16	100.0	428	4 AAU07219	AAu07219 Human asp
21	16	100.0	428	4 AAU07219	AAu07219 Human asp
22	16	100.0	428	4 AAU07219	AAu07219 Human asp
23	16	100.0	428	4 AAU07219	AAu07219 Human asp
24	16	100.0	428	4 AAU07219	AAu07219 Human asp
25	16	100.0	428	4 AAU07219	AAu07219 Human asp

26	16	100.0	434	4 AAU07220	AAu07220 Human asp
27	16	100.0	434	4 AAU07220	AAu07220 Human asp
28	16	100.0	434	4 AAU07220	AAu07220 Human asp
29	16	100.0	434	4 AAU07220	AAu07220 Human asp
30	16	100.0	434	4 AAU07220	AAu07220 Human asp
31	16	100.0	434	4 AAU07220	AAu07220 Human asp
32	16	100.0	434	4 AAU07220	AAu07220 Human asp
33	16	100.0	434	4 AAU07220	AAu07220 Human asp
34	16	100.0	434	4 AAU07220	AAu07220 Human asp
35	16	100.0	434	4 AAU07220	AAu07220 Human asp
36	16	100.0	434	4 AAU07220	AAu07220 Human asp
37	16	100.0	434	4 AAU07220	AAu07220 Human asp
38	16	100.0	434	4 AAU07220	AAu07220 Human asp
39	16	100.0	434	4 AAU07220	AAu07220 Human asp
40	16	100.0	434	4 AAU07220	AAu07220 Human asp
41	16	100.0	434	4 AAU07220	AAu07220 Human asp
42	16	100.0	434	4 AAU07220	AAu07220 Human asp
43	16	100.0	434	4 AAU07220	AAu07220 Human asp
44	16	100.0	434	4 AAU07220	AAu07220 Human asp
45	16	100.0	434	4 AAU07220	AAu07220 Human asp

#### ALIGNMENTS

RESULT 1	ADP83877	standard; peptide; 16 AA.
ID	ADP83877	standard; peptide; 16 AA.
AC	ADP83877	
XX	ADP83877	
DT	23-SEP-2004	(first entry)
DE	Human BAC1 22-37 prodomain amino acid sequence SEQ ID NO:2.	
XX	human; beta-site amyloid precursor protein cleaving enzyme 1;	
KW	beta-site APP cleaving enzyme 1; BAC1; BAC1 isoform A; chromosome 11;	
KW	prodomain; engineered cleavage site; protease domain; neuroprotective;	
KW	neurotropic; gene therapy; Alzheimer's disease; Down's syndrome.	
OS	Homo sapiens.	
XX	WO2004056962-A2.	
PN	08-JUL-2004.	
PD	02-DEC-2003; 2003WO-US038314.	
XX	04-DEC-2002; 2002US-0430984P.	
PR	(SUNE-) SUNEIS PHARM INC.	
PA	Ballinger M;	
XX	WPI; 2004-507703/48.	
PI	New polypeptides for producing homogeneously processed preparations of	
PT	beta site amyloid precursor protein-cleaving enzyme comprises a	
PT	prodomain, an engineered cleavage site and a protease domain.	
XX	Claim 1; SEQ ID NO 3; 40P; English.	
PS	The present invention describes a polypeptide (I) comprising in order	
CC	from the N-terminus to the C-terminus: (a) a prodomain comprising at	
CC	least 6 contiguous amino acids of the 16 amino acid sequence of SEQ ID	
CC	NO:3 (ADP83877), comprising residues 22-37 of SEQ ID NO:1 (ADP83876) which	
CC	is the longest isoform of human beta-site amyloid precursor protein (APP)	
CC	cleaving enzyme 1 (BAC1), isoform A; (b) an engineered cleavage site;	
CC	and (c) a protease domain. (I) is capable of being cleaved at the	
CC	engineered cleavage site, and so releases a free protease domain that has	
CC	BAC1 activity. Also described: (1) a nucleic acid sequence encoding (I);	
CC	(2) a vector for expression of (I); and (3) a host cell expressing (I).	
CC	(I) has neuroprotective and neurotropic activities, and can be used in gene	

CC therapy. (1) can be used for producing preparations of homogeneously  
CC processed BACE that may be used for e.g. studying or treating diseases  
CC such as Alzheimer's disease or Down's syndrome. The human BACE1 gene is  
CC located on chromosome 11, more specifically to 11q23.2-23.3. The present  
CC sequence represents the human BACE1 isoform A 22-37 prodomain amino acid  
CC sequence, which is used in the exemplification of the present invention.  
XX

Query Match 100.0%; Score 16; DB 8; Length 16;  
Best Local Similarity 100.0%; Pred. No. 7e-09;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
SQ Sequence 16 AA;  
QY 1 TONGIRLPRLRSGLGGA 16  
DB 1 TONGIRLPRLRSGLGGA 16

RESULT 2  
ADC81580  
ID ADC81580 standard; protein; 425 AA.  
XX  
AC ADC81580;  
XX  
DT 01-JAN-2004 (first entry)  
XX  
DE Beta-secretase zymogen (psbz) amino acid sequence SEQ ID NO:3.  
XX  
KM human; BACE; modification; Pro33lys; pro-enzyme.  
XX  
OS Unidentified.  
XX  
PN WO2003072733-A2.  
XX  
PD 04-SEP-2003.  
XX  
PF 21-FEB-2003; 2003WO-US005508.  
XX  
PR 21-FEB-2002; 2002US-0356651P.  
XX  
PA (PHAA) PHARMACIA & UPJOHN CO.  
XX  
PI Chou K, Howe JW;  
XX  
DR WPI; 2003-712719/67.  
XX  
PT BACE polypeptides having Pro33lys modification, useful in determining  
PT possible mutations, which will inhibit enzyme activity, and in  
PT determining potential active site for target molecules.  
XX  
PS Disclosure; Fig 3; 38pp; English.

CC The present invention describes an isolated polypeptide (1) comprising or  
CC consisting of a fully defined sequence of 432 amino acids (see ADC81561),  
CC and comprising human BACE having the modification Pro33lys. Also  
CC described: (1) a composition comprising an active human BACE enzyme  
CC comprising the pro-enzyme sequence of BACE having the modification  
CC Pro33lys; (2) an isolated polynucleotide comprising a sequence encoding  
CC (1); (3) an isolated polynucleotide consisting of a sequence encoding  
CC nucleotides 70-1365 of a 1355-bp sequence (see ADC81562); (4) an  
CC expression vector comprising the polynucleotide of (2); or a  
CC polynucleotide sequence encoding a Pro33lys-BACE polypeptide, where the  
CC expression vector can produce the Pro33lys-BACE polypeptide, where the  
CC in a compatible host cell, when cultured under conditions that allow  
CC production; (5) a recombinant host cell comprising the expression vector,  
CC and (6) producing a (active) Pro33lys-BACE polypeptide, the BACE  
CC polypeptide having Pro33lys modification may be used in determining  
CC possible mutations, which will inhibit enzyme activity, and in  
CC determining potential active site for target molecules. The vector  
CC comprising the BACE polynucleotide is useful for producing recombinant  
CC BACE polypeptides having Pro33lys modification. The present sequence  
CC represents a beta-secretase zymogen amino acid sequence, which is used in  
CC the exemplification of the present invention.

XX  
SQ Sequence 425 AA;

Query Match 100.0%; Score 16; DB 7; Length 425;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
QY 1 TONGIRLPRLRSGLGGA 16  
DB 1 TONGIRLPRLRSGLGGA 16

RESULT 3  
AAU07219  
ID AAU07219 standard; protein; 428 AA.  
XX  
AC AAU07219;  
XX  
DT 24-OCT-2001 (first entry)  
XX  
DE Human aspartyl protease 2b delatm (Huasp-2bdelatm).  
XX  
KM Human; aspartyl protease 1; Asp-1; neuroprotective;  
XX aspartyl protease 2; Asp2; amyloid protein precursor; App;  
XX beta-secretase; Alzheimer's disease; Huasp-2bdelatm.  
XX  
OS Homo sapiens.

Key Location/Qualifiers  
FH Misc-difference 1 /note= "Encoded by NNN"  
FT Misc-difference 2 /note= "Encoded by NNC"  
FT Misc-difference 61 /note= "Encoded by NNC"  
FT Misc-difference 62 /note= "Encoded by NNN"  
FT Misc-difference 121 /note= "Encoded by NNC"  
FT Misc-difference 122 /note= "Encoded by NNN"  
FT Misc-difference 181 /note= "Encoded by NNG"  
FT Misc-difference 182 /note= "Encoded by NNN"  
FT Misc-difference 241 /note= "Encoded by NNG"  
FT Misc-difference 242 /note= "Encoded by NNN"  
FT Misc-difference 301 /note= "Encoded by NNC"  
FT Misc-difference 302 /note= "Encoded by NNN"  
FT Misc-difference /note= "Encoded by NNT"  
PN WO200149097-A2.

PD 12-JUL-2001.  
XX  
PF 09-MAY-2001; 2001WO-IB000797.  
XX  
PR 09-MAY-2001; 2001WO-IB000797.  
XX  
PA (BIEN/) BIENKOWSKI M J.  
PA (GURN/) GURNEY M E.  
PA (HEIN/) HEINRIKSON R L.  
PA (PARO/) PARODI L A.  
XX (YANR/) YAN R.  
PI Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;  
XX WPI; 2001-502548/55.  
XX  
DR N-PSDB; AAS11732.  
XX



PT Novel purified polypeptide comprising fragment of mammalian aspartyl  
PT protease 2, lacking Asp2 transmembrane domain and retaining beta  
PT secretase activity of Asp2 useful for identifying inhibitors of Asp2  
PT activity.  
PS Claim 149, Page 167-168; 185pp; English.  
XX  
CC The invention relates to a novel purified polypeptide comprising a  
CC fragment of mammalian aspartyl protease 2 (Asp2) protein which lacks the  
CC Asp2 transmembrane domain and the Asp2 protein, and where the polypeptide  
CC and the fragment retain the beta-secretase activity of the mammalian Asp2  
CC protein. Also included is an isoform of amyloid precursor protein (APP)  
CC comprising the amino acid sequence of an APP or its fragment containing an  
CC APP cleavage site recognizable by a mammalian beta-secretase, and further  
CC comprising two lysine residues at the carboxyl terminus of the amino acid  
CC sequence of the mammalian APP or APP fragment. The polypeptides are used  
CC for assaying for modulators of beta-secretase activity; identifying  
CC agents that inhibit the APP processing activity of human Asp2 aspartyl  
CC protease (Hu-Asp2); identifying agents that modulate the activity of Asp2  
CC; and for reducing cellular production of amyloid beta (Abeta) from APP.  
CC Agents identified by the above methods are useful for treating  
CC Alzheimer's disease; and for identifying modulators of amyloid-beta  
CC (Abeta) peptide production, for use in designing therapeutics for the  
CC treatment or prevention of Alzheimer's disease. Probes and primers  
CC derived from Asp nucleic acid sequences are useful for detecting Hu-Asp  
CC nucleic acids in in vitro assays and in Northern and Southern blots. The  
CC present sequence represents the amino acid sequence of human Asp-2b delta  
CC TM construct which lacks the transmembrane domain. This construct was  
CC used for bacterial expression and purification of human Asp2b  
CC  
XX  
SO Sequence 428 AA;  
Query Match 100.0%; Score 16; DB 4; Length 428;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
QY 1 TQHGIRLPRLSGIGGA 16  
DB 22 TQHGIRLPRLSGIGGA 37  
RESULT 4  
AAE10646  
ID AAE10646 standard; protein; 428 AA.  
AC AAE10646;  
XX  
XX 10-DEC-2001 (first entry)  
DT Human-Asp 2(b) protein lacking transmembrane domain.  
DE Human-Asp 2(b) protein lacking transmembrane domain.  
XX  
XX Human; aspartyl protease 2b; Asp2b; amyloid precursor protein; APP;  
KW Alzheimer's disease; AD; dementia; neurofibrillary tangle; gliosis;  
KW amyloid plaque; neuronal loss; proteolytic; neurotropic; neuroprotective.  
XX  
XX Homo sapiens.  
OS Synthetic.  
XX  
XX GB2357767-A.  
PN  
XX  
PD 04-JUL-2001.  
XX  
PF 22-SEP-2000; 2000GB-00023315.  
XX  
XX 23-SEP-1999; 99US-00404133.  
PR 23-SEP-1999; 99US-0155493P.  
PR 23-SEP-1999; 99WO-US020881.  
PR 13-OCT-1999; 99US-00416901.  
PR 06-DEC-1999; 99US-0169232P.  
XX  
XX (PHAA ) PHARMACIA & UPJOHN CO.  
PA  
XX  
PI Bienkowski MJ, Gurney M;

XX  
XX WPI; 2001-444208/48.  
DR N-PSDB; AAD17895.  
XX  
XX Polypeptide comprising fragments of human aspartyl protease with amyloid  
PT precursor protein processing activity and alpha-secretase activity, for  
PT identifying modulators useful in treating Alzheimer's disease.  
XX  
XX Example 10; Page 138-139; 187pp; English.  
PS  
XX  
CC The patent discloses human aspartyl protease 1 (hu-Asp1) or modified Asp1  
CC proteins which lack transmembrane domain or amino terminal domain or  
CC cytoplasmic domain and retain alpha-secretase activity and amyloid  
CC protein precursor (APP) processing activity. The proteins of the  
CC invention are useful for assaying hu-Asp1 alpha-secretase activity, which  
CC in turn is useful for identifying modulators of hu-Asp1 alpha-secretase  
CC activity, where modulators that increase hu-Asp1 alpha-secretase activity  
CC are useful for treating Alzheimer's disease (AD) which causes progressive  
CC dementia with consequent formation of amyloid plaques, neurofibrillary  
CC tangles, gliosis and neuronal loss. Hu-Asp1 protease substrate is useful  
CC for assaying hu-Asp1 proteolytic activity, by contacting hu-Asp1 protein  
CC with the substrate under acidic conditions and determining the level of  
CC hu-Asp1 proteolytic activity. The present sequence is human Asp 2(b)  
CC protein lacking a transmembrane (TM) domain. This sequence is generated  
CC by the deletion of the C-terminal TM domain and intracellular domains of  
CC human Asp 2(b) protein  
XX  
SO Sequence 428 AA;  
Query Match 100.0%; Score 16; DB 4; Length 428;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
QY 1 TQHGIRLPRLSGIGGA 16  
DB 22 TQHGIRLPRLSGIGGA 37  
RESULT 5  
AAE06891  
ID AAE06891 standard; protein; 428 AA.  
AC AAE06891;  
XX  
XX 23-OCT-2001 (first entry)  
DT Human-Asp2(b) deltaTM protein.  
DE  
XX  
XX Human; aspartyl protease 2b; Asp 2b; beta-amyloid precursor protein; APP;  
KW beta-secretase; Alzheimer's disease; dementia; amyloid plaque; gliosis;  
KW neurofibrillary tangle; neuronal loss; amyloid-beta peptide; neurotropic;  
KW neuroprotective; antisense therapy; Asp2(b) deltaTM protein;  
KW gene therapy.  
XX  
XX Homo sapiens.  
OS Synthetic.  
XX  
XX WO200150829-A2.  
PN  
XX  
PD 19-JUL-2001.  
XX  
PF 09-MAY-2001; 2001WO-1B000799.  
XX  
XX 09-MAY-2001; 2001WO-1B000799.  
PR  
XX  
XX (BIEN/) BIENKOWSKI M J.  
PA (GURN/) GURNEY M B.  
PA (HEIN/) HEINRIKSON R L.  
PA (PARO/) PARODI L A.  
PA (YANR/) YAN R.  
XX  
XX Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;

DR WPI; 2001-483072/52.  
DR N-PSDB; AAD13276.

PT Novel purified polypeptide comprising fragment of mammalian aspartyl  
PT protease 2, lacking Asp2 transmembrane domain and retaining beta  
PT secretase activity of Asp2 useful for identifying inhibitors of Asp2  
PT activity.

PS Claim 149; Page 167-168; 185pp; English.

XX  
XX The invention relates to human aspartyl proteases (Hu-Asp), beta-amyloid  
CC precursor protein (APP) isoforms and their corresponding DNA molecules.  
CC Human aspartyl proteases can act as beta-secretase proteases useful for  
CC treating Alzheimer's disease. APP isoforms are useful for identifying  
CC modulators of amyloid-beta peptide production, for use in designing  
CC therapeutics for the treatment and prevention of Alzheimer's disease,  
CC dementia, formation of amyloid plaques, neurofibrillary tangles, gliosis  
CC and neuronal loss. APP isoforms are also used in methods for identifying  
CC inhibitors and modulators of human Asp2 activity. The invention relates  
CC to a method for identifying agents that modulate the activity of human  
CC aspartyl protease Asp2. Amyloid-beta peptides obtained from APP are used  
CC as a means to screen in cellular assays for the inhibitors of beta- and  
CC gamma-secretase. Hu-Asp DNA fragments are useful as probes or primers in  
CC polymerase chain reactions (PCR). The probes are useful for detecting Hu-  
CC Asp nucleic acids in in vitro assays and in Northern and Southern blots.  
CC The present sequence is Human aspartyl protease 2b (Hu-Asp2b) deltaTM  
CC protein which is obtained by the deletion of C-terminal transmembrane and  
CC intracellular domains of Hu-Asp2b. Human Asp2b has beta-secretase  
CC activity

SO Sequence 428 AA;

Query Match 100.0%; Score 16; DB 4; Length 428;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Oy 1 TOHGIRLPRLSGGGA 16  
|||  
Db 22 TOHGIRLPRLSGGGA 37

RESULT 6  
AAE02598  
ID AAE02598 standard; protein; 428 AA.  
XX  
AC AAE02598;

DT 10-AUG-2001 (first entry)  
XX

DE Human aspartyl protease 2 (b) delta TM protein.

XX Human; alpha-secretase; amyloid precursor protein; APP; therapy;  
KW Alzheimer's disease; antialzheimer's; aspartyl protease 2; Asp 2;  
KW beta-secretase; chromosome 11q23.3-24.1; mutant; mutein.

OS Homo sapiens.  
OS Synthetic.

PN WO200123533-A2.

PD 05-APR-2001.

PF 22-SEP-2000; 2000WO-US026080.

PR 23-SEP-1999; 99US-0155493P.

PR 23-SEP-1999; 99WO-US020881.

PR 13-OCT-1999; 99US-00416901.

PR 06-DEC-1999; 99US-0169232P.

XX (PHAA ) PHARMACIA & UPJOHN CO.

PA Gurney M, Bienkowski MJ;  
XX

DR WPI; 2001-290516/30.

DR N-PSDB; AAD06768.

XX  
XX Enzymes that cleave the alpha-secretase site of the amyloid precursor  
PT protein, useful for the treatment of Alzheimer's disease.

PS Example 10; Page 166-167; 189pp; English.

XX  
XX The present invention relates to enzymes for cleaving the alpha-  
CC secretase site of the amyloid precursor protein (APP) and methods of  
CC identifying those enzymes. The methods may be used to identify enzymes  
CC that may be used to cleave the alpha-secretase cleavage site of the APP  
CC protein. The enzymes may be used to treat or modulate the progress of  
CC Alzheimer's disease. The present sequence is human aspartyl protease 2  
CC (Asp 2) (b) delta TM protein. The Asp 2 gene is located on chromosome  
CC 11q23.3-24.1. The Asp 2 has beta-secretase protease activity

SO Sequence 428 AA;

Query Match 100.0%; Score 16; DB 4; Length 428;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Oy 1 TOHGIRLPRLSGGGA 16  
|||  
Db 22 TOHGIRLPRLSGGGA 37

RESULT 7  
AAU06620  
ID AAU06620 standard; protein; 428 AA.  
XX  
AC AAU06620;

DT 24-OCT-2001 (first entry)  
XX

DE Human-pro-Asp 2 (b) delta TM.

XX Human; Aspartyl protease; beta-secretase; neurotropic; ASP2;  
KW neuroprotective; amyloid protein precursor; APP; Alzheimer's disease;  
KW amyloid-beta; Abeta; Human-pro-Asp 2 (b) delta TM; mutant; mutein.

OS Homo sapiens.  
OS Synthetic.

PN WO200149098-A2.

PD 12-JUL-2001.

PF 09-MAY-2001; 2001WO-IB000798.

PR 09-MAY-2001; 2001WO-IB000798.

XX (BIEN/) BIENKOWSKI M J.

PA (GURN/) GURNEY M E.

PA (HEIN/) HEINRIKSON R L.

PA (PARO/) PARODI L A.

PA (YANR/) YAN R.

PI Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;  
XX

DR WPI; 2001-502549/55.

XX  
XX Novel purified polypeptide comprising fragment of mammalian aspartyl  
PT protease 2, lacking Asp2 transmembrane domain and retaining beta  
PT secretase activity of Asp2 useful for identifying inhibitors of Asp2  
PT activity.

PS Claim 149; Page 167-168; 185pp; English.

XX The invention relates to a purified polypeptide comprising a fragment of  
CC mammalian aspartyl protease (Asp)2 protein which lacks the Asp2  
CC transmembrane domain and the Asp2 protein, and where the polypeptide and

CC the fragment retain the beta-secretase activity of the mammalian Asp2  
CC protein. The invention also details polynucleotides for the Asp proteins  
CC and vectors expressing them, and a polypeptide (isoform of amyloid  
CC protein precursor (APP)) comprising the amino acid sequence of an APP or  
CC its fragment containing an APP cleavage site recognizable by a mammalian  
CC beta-secretase, and further comprising two lysine residues at the  
CC carboxyl terminus of the amino acid sequence of the mammalian APP or APP  
CC fragment. Also included in the invention are methods of identifying  
CC modulators or inhibitors of Asp2. Modulators and inhibitors of Asp2 are  
CC useful for treating Alzheimer's disease. APP is useful in methods for  
CC identifying inhibitors or modulators of human Asp2 activity and amyloid-  
CC beta (Abeta) peptide production. APP is also useful in designing  
CC therapeutics for the treatment or prevention of Alzheimer's disease. APP  
CC comprising the APP-Sw-beta-secretase peptide sequence (NLDA), which is  
CC associated with increased levels of Abeta processing is useful in assays  
CC relating the Alzheimer's research. The expression vector is useful for  
CC recombinantly expressing APP. Nucleic acids that hybridize to APP  
CC oligonucleotides are useful as probes or primers. The probes are useful  
CC for detecting Hu-Asp nucleic acids in *in vitro* assays and in Northern and  
CC Southern blots. The present sequence is Human-pro- Asp 2(b) delta TM  
CC protein, which lacks the C-terminal transmembrane domain  
CC  
XX  
SQ Sequence 428 AA;

Query Match 100.0%; Score 16; DB 4; Length 428;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 TQHGIRLPRLPSGLGGA 16  
|||  
22 TQHGIRLPRLPSGLGGA 37

RESULT 8  
AB878607  
ID AB878607 standard; protein; 428 AA.

AC AB878607;

DT 16-JUN-2002 (first entry)

DE Human Asp-2(b) deltatm protein sequence SEQ ID NO:51.

KM Human; Asp-1; Asp-2; aspartyl protease; Alzheimer's disease; proteolytic;  
XX chromosome 11q23.3-24.1.

OS Homo sapiens.

KW GB2367060-A.

PN GB2367060-A.

XX 27-MAR-2002.

PD 27-MAR-2002.

PF 29-OCT-2001; 2001GB-00025934.

XX 23-SEP-1999; 99US-00404133.

PR 23-SEP-1999; 99US-0155493P.

PR 23-SEP-1999; 99WO-US020881.

PR 13-OCT-1999; 99US-00416901.

PR 06-DEC-1999; 99US-0169232P.

PR 22-SEP-2000; 2000GB-00023315.

XX (PHAA ) PHARMACIA & UPJOHN CO.

PA (PHAA ) PHARMACIA & UPJOHN CO.

PI Bienkowski MJ, Gurney M;

XX MPI: 2002-397167/43.

DR N-PSDB; ABL52487.

XX Human aspartyl protease 1 substrates useful in assays to detect aspartyl

PT protease activity, e.g. for the diagnosis of Alzheimer's disease.

XX Example 10; Page 138-139; 182pp; English.

CC The present invention describes a human aspartyl protease 1 (hu-Asp1)  
CC substrate (I) which comprises a peptide of no more than 50 amino acids,  
CC and which comprises the 8 amino acid sequence Gly-Leu-Ala-Leu-Ala-Leu-  
CC Glu-Pro. Also described are: (1) a method (II) for assaying hu-Asp1  
CC proteolytic activity, comprising: (a) contacting a hu-Asp1 protein with  
CC (I) under acidic conditions; and (b) determining the level of hu-Asp1  
CC proteolytic activity; (2) a purified polynucleotide (III) comprising a  
CC nucleotide sequence that hybridizes under stringent conditions to the non  
CC coding strand complementary to a defined 1804 nucleotide sequence (see  
CC AB52456) where the nucleotide sequence encodes a polypeptide having Asp1  
CC proteolytic activity and lacks nucleotides encoding a transmembrane  
CC domain; (3) a purified polynucleotide (III') comprising a sequence that  
CC hybridizes under stringent conditions to (III) (the nucleotide sequence  
CC encodes a polypeptide further lacking a pro-peptide domain corresponding  
CC to amino acids 23-62 of hu-Asp1 (see AB878589)); (4) a vector (IV)  
CC comprising (III) or (III') and (5) a host cell (V) transformed or  
CC transfected with (III), (III') and/or (IV). The hu-Asp1 protease  
CC substrate (I) may be used as an enzyme substrate in assays to detect  
CC aspartyl protease activity, (II) and therefore diagnose diseases  
CC associated with aberrant hu-Asp1 expression and activity such as  
CC Alzheimer's disease. Hu-Asp1 has been localised to chromosome 21, while  
CC hu-Asp2 has been localised to chromosome 11q23.3-24.1. The present  
CC sequence represents human Asp-2(b) deltatm, which is given in an example  
CC from the present invention  
CC  
XX  
SQ Sequence 428 AA;

Query Match 100.0%; Score 16; DB 5; Length 428;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 TQHGIRLPRLPSGLGGA 16  
|||  
22 TQHGIRLPRLPSGLGGA 37

RESULT 9  
ADJ94363  
ID ADJ94363 standard; protein; 428 AA.

AC ADJ94363;

DT 03-JUN-2004 (first entry)

DE Human-pro-Asp-2(b) deltatm.

XX Human; enzyme; aspartyl protease; Asp-1; Asp-2(a); Asp-2(b);

XX beta secretase; amyloid protein precursor; APP; Alzheimer's disease;

XX neurotropic; neuroprotective; amyloid beta; mutant; mutain.

OS Homo sapiens.

OS Synthetic.

XX US6706485-B1.

PN US6706485-B1.

XX 16-MAR-2004.

PD 12-APR-2000; 2000US-00548376.

XX 24-SEP-1998; 98US-0101594P.

PR 23-SEP-1999; 99US-00404133.

PR 23-SEP-1999; 99US-0155493P.

PR 23-SEP-1999; 99WO-US020881.

PR 13-OCT-1999; 99US-00416901.

XX (PHAA ) PHARMACIA & UPJOHN CO.

PA (PHAA ) PHARMACIA & UPJOHN CO.

PI Gurney ME, Bienkowski MJ, Heinrikson RL, Parodi LA, Yan R;

XX MPI: 2004-236722/22.

DR N-PSDB; ADJ94362.

XX Identifying agents that modulate activity of Asp2 aspartyl protease

PT

PT useful for treating or preventing Alzheimer's disease involves comparing  
PT APP processing activity of protease in presence and absence of test  
PT agent.

XX Example 10; SEQ ID NO 51; 109pp; English.

CC The invention relates to identifying agents that modulate activity of  
CC Asp2 (e.g., a beta-secretase, e.g., human Asp-2(b) appearing as ID 6,  
CC encoded by ID 5) aspartyl protease, involves contacting Asp2 with amyloid  
CC precursor protein (APP) in the presence and absence of a test agent,  
CC where Asp2 is a recombinant polypeptide and processes APP into amyloid  
CC beta, determining APP processing activity of Asp2 in presence and absence  
CC of the test agent, and comparing the activities to identify agents that  
CC modulate the activity of Asp2. Also disclosed are the cDNA and proteins  
CC for human Asp-1 and Asp-2(a), mouse Asp-2(b), a vector comprising the  
CC nucleic acid encoding Hu-Asp2 protease sequence, a host cell comprising  
CC the vector and the method of producing Hu-Asp polypeptide, an isolated  
CC antibody that specifically binds to Hu-Asp polypeptides, identifying a  
CC cell that can be used to screen for inhibitors of beta secretase  
CC activity, novel isoforms of amyloid protein precursor (APP), where the  
CC last 2 carboxy terminus amino acids of that isoform are both lysine  
CC residues (e.g., those designated APP695-KK or carrying the Swedish  
CC mutation where KM at 595-596 is mutated to NL, designated e.g., APP695-SW  
CC or APP695-SW-KK, or a V to F mutation at 642, e.g., APP695-VF, all useful  
CC for assaying for beta secretase activity and screening for inhibitors of  
CC beta-secretase) and polynucleotides that encode the APP proteins. The  
CC method is useful for identifying agents that modulate the activity  
CC (amyloid precursor protein processing activity) of Asp2 aspartyl  
CC protease. Preferably, the method is useful for identifying agents that  
CC inhibit Asp2 aspartyl protease activity. The inhibitors of amyloid  
CC precursor protein processing, are useful for treating or preventing  
CC Alzheimer's disease. The present sequence represents an aspartyl protease  
CC mutant construct (e.g., lacking a transmembrane domain and/or including a  
CC caspase cleavage site) used to investigate the cleavage activity of Asp2  
CC proteins.

XX SQ Sequence 428 AA;

Query Match 100.0%; Score 16; DB 8; Length 428;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Oy 1 TQHGIRLPRLSGIGCA 16  
|||  
Db 22 TQHGIRLPRLSGIGCA 37

RESULT 10

AD050459 ID AD050459 standard; protein; 428 AA.

XX AC AD050459;

XX DT 29-JUL-2004 (first entry)

XX DE Human Asp2(b) deltatm mutant protein.

XX KM Aspartyl protease; Asp, beta secretase; amyloid precursor protein; APP;

XX KM Alzheimer's disease; gene therapy; human; mutant; mulein.

XX OS Homo sapiens.

XX OS Synthetic.

XX PN US6737510-B1.

XX PD 18-MAY-2004.

XX PF 12-APR-2000; 2000US-00548373.

XX PR 24-SEP-1998; 98US-0101594P.  
XX PR 23-SEP-1999; 99US-00404133.  
XX PR 23-SEP-1999; 99US-0155493P.  
XX PR 23-SEP-1999; 99WO-US020881.

PR 13-OCT-1999; 99US-00416901.

XX PA (PHAA ) PHARMACIA & UPJOHN CO.

XX PI Gurney ME, Bienkowiaki MJ, Heinrikson RL, Parodi LA, Yan R;

XX DR WPI; 2004-387112/36.

XX DR N-PSDB; AD050458.

PT New Asp2 aspartyl protease protein comprising tripeptides DTG and DSG

PT involved in processing amyloid precursor protein into amyloid beta,

PT useful in preparing a composition for treating or preventing Alzheimer's

PT disease.

PS Example 10; SEQ ID NO 51; 108pp; English.

XX The invention relates to a method for identifying an agent that decreases

CC the protease activity of the aspartyl protease (Asp) polypeptide. It also

CC provides enzyme and enzymatic procedures for cleaving the beta secretase

CC cleavage site of the amyloid precursor protein (APP). The invention is

CC useful in preparing a composition for treating or preventing Alzheimer's

CC disease. It is also useful in gene therapy. The present sequence is human

CC Asp2(b) mutant protein. This sequence is used to illustrate the method of

XX the invention.

XX SQ Sequence 428 AA;

Query Match 100.0%; Score 16; DB 8; Length 428;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Oy 1 TQHGIRLPRLSGIGCA 16  
|||  
Db 22 TQHGIRLPRLSGIGCA 37

RESULT 11

AD75372 ID AD75372 standard; protein; 428 AA.

XX AC AD75372;

XX DT 18-NOV-2004 (first entry)

XX DE Human Asp2(b) deltatm mutant protein.

XX KM Aspartyl protease; Asp; amyloid precursor protein; APP; amyloid beta;

XX KM chromosome identification; Alzheimer's disease; human; mutant.

XX OS Homo sapiens.

XX OS Synthetic.

XX PN US2004166507-A1.

XX PD 26-AUG-2004.

XX PF 29-AUG-2003; 2003US-00652045.

XX PR 24-SEP-1998; 98US-0101594P.  
XX PR 23-SEP-1999; 99US-00404133.  
XX PR 23-SEP-1999; 99US-0155493P.  
XX PR 13-OCT-1999; 99US-00416901.

XX PA (GURN/) GURNEY M E.  
XX PA (BIEN/) BIENKOWAKI M J.  
XX PA (HEIN/) HEINRIKSON R L.  
XX PA (PARO/) PARODI L A.  
XX PA (YANR/) YAN R.  
XX PT Gurney ME, Bienkowiaki MJ, Heinrikson RL, Parodi LA, Yan R;  
XX DR WPI; 2004-624916/60.  
XX DR N-PSDB; AD75371.

XX Novel purified/isolated polynucleotide encoding polypeptide having  
PT aspartyl protease activity involved in processing amyloid precursor  
PT protein into amyloid beta, useful in identifying agent decreasing  
PT activity of aspartyl protease.  
XX  
PS Example 10; SEQ ID NO 51; 107pp; English.  
XX The invention relates to nucleic acid sequences encoding aspartyl  
CC protease (Asp) polypeptides having aspartyl protease activity involved in  
CC processing amyloid precursor protein (APP) into amyloid beta. The  
CC invention also relates to a method for identifying an agent that  
CC decreases the protease activity of the Asp. Asp DNA is useful in  
CC chromosome identification as they can hybridise with a specific location  
CC on a human chromosome and in identifying the relationship between genes  
CC and diseases (particular gene responsible for causing diseases). It is  
CC also useful for identifying candidates to modulate the progression of  
CC Alzheimer's disease. Asp is useful in raising antibodies that are useful  
CC in diagnostic assay for detecting Hu-Asp polypeptide expression. The  
CC present sequence is the human Asp2(b)deltaTM mutant protein. This  
CC sequence is used to illustrate the method of the invention.  
XX  
SQ Sequence 428 AA;  
Query Match 100.0%; Score 16; DB 8; Length 428;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
OY 1 TQHGIRLPLRSLGGA 16  
DB 22 TQHGIRLPLRSLGGA 37  
RESULT 12  
ADC81561  
ID ADC81561 standard; protein; 432 AA.  
XX  
AC ADC81561;  
XX  
DT 01-JAN-2004 (first entry)  
XX  
DE Mature BACE p33K amino acid sequence SEQ ID NO:2.  
XX  
KM human; BACE; modification; Pro33lys; pro-enzyme.  
XX  
OS Synthetic.  
OS Homo sapiens.  
OS  
PN WO2003072733-A2.  
XX  
PD 04-SEP-2003.  
XX  
PF 21-FEB-2003; 2003WO-US005508.  
XX  
PR 21-FEB-2002; 2002US-0358651P.  
XX  
PA (PHAA ) PHARMACIA & UPJOHN CO.  
XX  
PI Chou K, Howe JW;  
XX  
DR MPI: 2003-712719/67.  
DR N-PSDB; ADC81562.  
XX  
PT BACE polypeptides having Pro33lys modification, useful in determining  
PT possible mutations, which will inhibit enzyme activity, and in  
PT determining potential active site for target molecules.  
XX  
PS Claim 10; SEQ ID NO 2; 38pp; English.  
XX  
CC The present invention describes an isolated polypeptide (1) comprising or  
CC consisting of a fully defined sequence of 432 amino acids (see ADC81561),  
CC and comprising human BACE having the modification Pro33lys. Also  
CC described: (1) a composition comprising an active human BACE enzyme

CC comprising the pro-enzyme sequence of BACE having the modification  
CC Pro33lys; (2) an isolated polynucleotide comprising a sequence encoding  
CC (1); (3) an isolated polynucleotide consisting of or comprising of  
CC nucleotides 70-1165 of a 1355-bp sequence (see ADC81562); (4) an  
CC expression vector comprising the polynucleotide of (2), or a  
CC polynucleotide sequence encoding a Pro33lys-BACE polypeptide, where the  
CC expression vector can produce the Pro33lys-BACE polypeptide when present  
CC in a compatible host cell, when cultured under conditions that allow  
CC production; (5) a recombinant host cell comprising the expression vector;  
CC and (6) producing a (active) Pro33lys-BACE polypeptide. The BACE  
CC polypeptide having Pro33lys modification may be used in determining  
CC possible mutations, which will inhibit enzyme activity, and in  
CC determining potential active site for target molecules. The vector  
CC comprising the BACE polynucleotide is useful for producing recombinant  
CC BACE polypeptides having Pro33lys modification. The present sequence  
CC represents the mature recombinant BACE p33K amino acid sequence used in  
CC the exemplification of the present invention.  
XX  
SQ Sequence 432 AA;  
Query Match 100.0%; Score 16; DB 7; Length 432;  
Best Local Similarity 100.0%; Pred. No. 1.1e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
OY 1 TQHGIRLPLRSLGGA 16  
DB 1 TQHGIRLPLRSLGGA 16  
RESULT 13  
AAV88433  
ID AAV88433 standard; protein; 433 AA.  
XX  
AC AAV88433;  
XX  
DT 12-SEP-2003 (revised)  
DT 06-AUG-2003 (revised)  
DT 03-AUG-2000 (first entry)  
XX  
DE Human-pro-Asp-2(a)-deltaTM amino acid sequence.  
XX  
KM Aspartyl protease; aspartase; amyloid precursor protein; APP; Asp 2;  
XX Alzheimer's disease; beta secretase site; human-pro-Asp-2(a)-deltaTM.  
XX  
OS Homo sapiens.  
OS Enterobacteria phage T7.  
OS Chimeric.  
OS  
PN WO200017369-A2.  
XX  
PD 30-MAR-2000.  
XX  
PF 23-SEP-1999; 99WO-US020881.  
XX  
PR 24-SEP-1998; 98US-0101594P.  
XX  
PA (PHAA ) PHARMACIA & UPJOHN CO.  
XX  
PI Gurney ME, Bienkowski MJ, Heinrichson RL, Parodi LA, Van R;  
XX  
DR MPI: 2000-303209/26.  
DR N-PSDB; AAA15670.  
XX  
PT New enzyme designated human aspartase useful in research into Alzheimer's  
PT disease is capable of cleaving amyloid protein precursor at the beta  
PT secretase site to produce amyloid beta peptide.  
XX  
PS Example 9; Fig 8; 183pp; English.  
XX  
CC This sequence represents a modified version of the human aspartase 2  
CC (Asp2) amino acid sequence. The sequence is used in the bacterial  
CC expression of human Asp2L. The invention relates to a protease (e.g.  
CC Asp2) capable of cleaving the beta secretase site of amyloid precursor

CC protein (App). The protease contains a sequence encoding the amino acid  
CC sequence DTG and a sequence encoding DSG or DRG separated by 100-300  
CC amino acids. When mutated the App gene causes an autosomal dominant form  
CC of Alzheimer's disease. App localises to the cell surface membrane and  
CC have a single C-terminal transmembrane domain. Proteolytic processing of  
CC APP produces the amyloid beta protein, which is possibly very important  
CC in Alzheimer's disease. The invention includes a nucleotide sequence  
CC encoding the protease, a vector containing the nucleotide sequence, and a  
CC cell line comprising the vector. Methods for screening for inhibitors of  
CC beta secretase activity are also given in the invention. The human  
CC aspartase protein and nucleotide sequences and the methods for  
CC identifying inhibitors of the protease, are useful in the treatment of  
CC and research in to Alzheimer's disease. (Updated on 06-AUG-2003 to  
CC correct OS field.) (Updated on 12-SEP-2003 to standardise OS field)

CC Sequence 433 AA;

Query Match Best Local Similarity 100.0%; Score 16; DB 3; Length 433;

Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 1 TOHGIRLPLRSGIGCA 16  
Db 2 TOHGIRLPLRSGIGCA 17

RESULT 14

AAU07213 AAU07213 standard; protein; 433 AA.

AC AAU07213;

DT 11-SEP-2003 (revised)

DT 24-OCT-2001 (first entry)

DE T7-human aspartyl protease 2a delcatm (low GC).

XX Human; aspartyl protease 1; Asp-1; nootropic; neuroprotective;

KW aspartyl protease 2; Asp2; amyloid protein precursor; App;

KW beta-secretase; Alzheimer's disease.

OS Homo sapiens.

OS Enterobacteria phage T7.

XX WO200149097-A2.

XX 12-JUL-2001.

XX 09-MAY-2001; 2001WO-IB000797.

XX 09-MAY-2001; 2001WO-IB000797.

XX 09-MAY-2001; 2001WO-IB000797.

XX (BIEN/) BIENKOWSKI M J.

XX (GURNEY) GURNEY M E.

XX (HEIN/) HEINRIKSON R L.

XX (PARO/) PARODI L A.

XX (YANR/) YAN R.

XX Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;

XX WPI, 2001-502548/55.

XX N-PSDB; AAS11713.

XX Novel purified polypeptide comprising fragment of mammalian aspartyl

XX protease 2, lacking Asp2 transmembrane domain and retaining beta

XX secretase activity of Asp2 useful for identifying inhibitors of Asp2

XX activity.

XX Example 9; Fig 8; 185pp; English.

XX The invention relates to a novel purified polypeptide comprising a

XX fragment of mammalian aspartyl protease 2 (Asp2) protein which lacks the

XX Asp2 transmembrane domain and the Asp2 protein, and where the polypeptide

CC and the fragment retain the beta-secretase activity of the mammalian Asp2  
CC protein. Also included is an isoform of amyloid protein precursor (APP)  
CC comprising the amino acid sequence of a App or its fragment containing an  
CC APP cleavage site recognisable by a mammalian beta-secretase, and further  
CC comprising two lysine residues at the carboxyl terminus of the amino acid  
CC sequence of the mammalian App or App fragment. The polypeptides are used  
CC for assaying for modulators of beta-secretase activity; identifying  
CC agents that inhibit the APP processing activity of human Asp2 aspartyl  
CC protease (Hu-Asp2); identifying agents that modulate the activity of Asp2  
CC ; and for reducing cellular production of amyloid beta (Abeta) from APP.  
CC Alzheimer's disease, for identifying modulators of amyloid-beta (Abeta)  
CC peptide production, and for use in designing therapeutics for the  
CC treatment or prevention of Alzheimer's disease. Probes and primers  
CC derived from Asp nucleic acid sequences are useful for detecting Hu-Asp  
CC nucleic acids in in vitro assays and in Northern and Southern blots. The  
CC present sequence represents the amino acid sequence of T7-human Asp-2a  
CC delta TM (low GC) construct which has a T7 tag, has the GC content of the  
CC 5' sequence reduced by site-directed mutagenesis, and lacks the  
CC transmembrane domain. This construct was used for bacterial expression  
CC and purification of human Asp2a. (Updated on 11-SEP-2003 to standardise  
CC OS field)

CC Sequence 433 AA;

Query Match Best Local Similarity 100.0%; Score 16; DB 4; Length 433;

Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 1 TOHGIRLPLRSGIGCA 16  
Db 2 TOHGIRLPLRSGIGCA 17

RESULT 15

AAE10640 AAE10640 standard; protein; 433 AA.

AC AAE10640;

DT 10-DEC-2001 (first entry)

DE Human-pro-Asp 2(a) protein lacking TM domain.

XX Human; aspartyl protease 1; Asp1; amyloid precursor protein; App;

KW Alzheimer's disease; AD; dementia; neurofibrillary tangle; gliosis;

KW amyloid plaque; neuronal loss; proteolytic; nootropic; neuroprotective;

KW Human-pro-Asp 2(a) protein.

XX Homo sapiens.

XX Synthetic.

XX GB2357767-A.

XX 04-JUL-2001.

XX 22-SEP-2000; 2000GB-00023315.

XX 23-SEP-1999; 99US-00404133.

XX 23-SEP-1999; 99US-0155493P.

XX 23-SEP-1999; 99WO-US020881.

XX 13-OCT-1999; 99US-00416901.

XX 06-DEC-1999; 99US-0169232P.

XX (PHAA ) PHARMACIA & UPJOHN CO.

XX Bienkowski MJ, Gurney M;

XX WPI, 2001-444208/48.

XX N-PSDB; AAD17876.

XX Polypeptide comprising fragments of human aspartyl protease with amyloid

XX precursor protein processing activity and alpha-secretase activity, for



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## OM protein - protein search, using sw model

Run on: July 26, 2005, 16:31:35 ; Search time 153 Seconds  
(without alignments)  
40.679 Million cell updates/sec

Title: US-10-726-967A-3  
Perfect score: 16  
Sequence: 1 TOHGIRLPRLRSGLGGA 16

Scoring table: OLIGO  
Gapop 60.0 , Gapext 60.0

Searched: 1741741 seqs, 38892284 residues

Word size : 0

Total number of hits satisfying chosen parameters: 1741741

Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Listing first 45 summaries

Database :

Published Applications AA:\*

- 1: /cgn2\_6/ptodata/2/pubpaa/US07\_PUBCOMB.pep:\*
- 2: /cgn2\_6/ptodata/2/pubpaa/PCT\_NEW\_PUB.pep:\*
- 3: /cgn2\_6/ptodata/2/pubpaa/US06\_NEW\_PUB.pep:\*
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- 6: /cgn2\_6/ptodata/2/pubpaa/PCTUS\_PUBCOMB.pep:\*
- 7: /cgn2\_6/ptodata/2/pubpaa/US08\_NEW\_PUB.pep:\*
- 8: /cgn2\_6/ptodata/2/pubpaa/US08\_PUBCOMB.pep:\*
- 9: /cgn2\_6/ptodata/2/pubpaa/US09\_PUBCOMB.pep:\*
- 10: /cgn2\_6/ptodata/2/pubpaa/US09\_PUBCOMB.pep:\*
- 11: /cgn2\_6/ptodata/2/pubpaa/US09C\_PUBCOMB.pep:\*
- 12: /cgn2\_6/ptodata/2/pubpaa/US09C\_PUBCOMB.pep:\*
- 13: /cgn2\_6/ptodata/2/pubpaa/US10\_PUBCOMB.pep:\*
- 14: /cgn2\_6/ptodata/2/pubpaa/US10\_PUBCOMB.pep:\*
- 15: /cgn2\_6/ptodata/2/pubpaa/US10C\_PUBCOMB.pep:\*
- 16: /cgn2\_6/ptodata/2/pubpaa/US10D\_PUBCOMB.pep:\*
- 17: /cgn2\_6/ptodata/2/pubpaa/US10E\_PUBCOMB.pep:\*
- 18: /cgn2\_6/ptodata/2/pubpaa/US10E\_PUBCOMB.pep:\*
- 19: /cgn2\_6/ptodata/2/pubpaa/US11\_NEW\_PUB.pep:\*
- 20: /cgn2\_6/ptodata/2/pubpaa/US11\_NEW\_PUB.pep:\*
- 21: /cgn2\_6/ptodata/2/pubpaa/US60\_NEW\_PUB.pep:\*
- 22: /cgn2\_6/ptodata/2/pubpaa/US60\_PUBCOMB.pep:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	16	100.0	16	17	US-10-726-967A-3
2	16	100.0	425	15	US-10-372-473-3
3	16	100.0	428	9	US-09-794-927-51
4	16	100.0	428	9	US-09-795-847-51
5	16	100.0	428	9	US-09-794-743-51
6	16	100.0	428	9	US-09-794-748-51
7	16	100.0	428	9	US-09-794-925-51
8	16	100.0	428	9	US-09-681-442-51
9	16	100.0	428	10	US-09-869-414-51
10	16	100.0	428	10	US-09-548-366-51
11	16	100.0	428	15	US-10-652-927-51

12	16	100.0	428	15	US-10-652-830-51	Sequence 51, Appl
13	16	100.0	428	16	US-10-652-045-51	Sequence 51, Appl
14	16	100.0	428	16	US-10-476-935-51	Sequence 51, Appl
15	16	100.0	428	17	US-10-477-076-51	Sequence 51, Appl
16	16	100.0	432	15	US-10-372-473-2	Sequence 2, Appl
17	16	100.0	433	9	US-09-794-927-26	Sequence 26, Appl
18	16	100.0	433	9	US-09-795-847-26	Sequence 26, Appl
19	16	100.0	433	9	US-09-794-743-26	Sequence 26, Appl
20	16	100.0	433	9	US-09-794-748-26	Sequence 26, Appl
21	16	100.0	433	9	US-09-794-925-26	Sequence 26, Appl
22	16	100.0	433	9	US-09-681-442-26	Sequence 26, Appl
23	16	100.0	433	10	US-09-869-414-26	Sequence 26, Appl
24	16	100.0	433	10	US-09-548-366-26	Sequence 26, Appl
25	16	100.0	433	15	US-10-652-927-26	Sequence 26, Appl
26	16	100.0	433	15	US-10-652-830-26	Sequence 26, Appl
27	16	100.0	433	16	US-10-652-045-26	Sequence 26, Appl
28	16	100.0	433	16	US-10-476-935-26	Sequence 26, Appl
29	16	100.0	433	17	US-10-940-867-26	Sequence 26, Appl
30	16	100.0	433	17	US-10-726-967A-78	Sequence 78, Appl
31	16	100.0	433	17	US-10-726-967A-81	Sequence 81, Appl
32	16	100.0	433	17	US-10-726-967A-84	Sequence 84, Appl
33	16	100.0	433	17	US-10-477-076-26	Sequence 26, Appl
34	16	100.0	434	9	US-09-794-927-53	Sequence 53, Appl
35	16	100.0	434	9	US-09-795-847-53	Sequence 53, Appl
36	16	100.0	434	9	US-09-794-743-53	Sequence 53, Appl
37	16	100.0	434	9	US-09-794-748-53	Sequence 53, Appl
38	16	100.0	434	9	US-09-794-925-53	Sequence 53, Appl
39	16	100.0	434	9	US-09-681-442-53	Sequence 53, Appl
40	16	100.0	434	10	US-09-869-414-53	Sequence 53, Appl
41	16	100.0	434	10	US-09-548-366-53	Sequence 53, Appl
42	16	100.0	434	15	US-10-652-927-53	Sequence 53, Appl
43	16	100.0	434	15	US-10-652-830-53	Sequence 53, Appl
44	16	100.0	434	16	US-10-652-045-53	Sequence 53, Appl
45	16	100.0	434	16	US-10-476-935-53	Sequence 53, Appl

## ALIGNMENTS

RESULT 1  
US-10-726-967A-3  
; Sequence 3, Application US/10726967A  
; Publication No. US20050074456A1  
GENERAL INFORMATION:  
; APPLICANT: Ballinger, Marcus  
; TITLE OF INVENTION: Constructs for Homogenously Processed Preparations of Beta Site  
; FILE REFERENCE: 2004345-0021  
; CURRENT APPLICATION NUMBER: US/10/726,967A  
; CURRENT FILING DATE: 2003-12-02  
; NUMBER OF SEQ ID NOS: 110  
; SOFTWARE: PatentIn version 3.2  
; SEQ ID NO 3  
; LENGTH: 16  
; TYPE: PRT  
; ORGANISM: Artificial  
; FEATURE:  
; OTHER INFORMATION: Residues 22-37 of human BACE1 preprosequence  
US-10-726-967A-3

Query Match 100.0%; Score 16; DB 17; Length 16;  
Best Local Similarity 100.0%; Pred. No. 1,7e+08;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
QY 1 TOHGIRLPRLRSGLGGA 16  
DB 1 TOHGIRLPRLRSGLGGA 16  
RESULT 2  
US-10-372-473-3  
; Sequence 3, Application US/10372473  
; Publication No. US20040005691A1



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; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 51
; LENGTH: 428
; TYPE: PRT
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
; OTHER INFORMATION: delta TM
US-09-794-743-51

Query Match          100.0%; Score 16; DB 9; Length 428;
Best Local Similarity 100.0%; Pred. No. 2.6e-07;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 TOHGIRLPLRSGGGA 16
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        22 TOHGIRLPLRSGGGA 37

DB      22 TOHGIRLPLRSGGGA 37

RESULT 6
US-09-794-748-51
; Sequence 51, Application US/09794748
; Patent No. US20020037315A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrikson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND
; TITLE OF INVENTION: USES
; TITLE OF INVENTION: THEREFOR
; FILE REFERENCE: 28341/6280JL
; CURRENT FILING DATE: 2001-02-27
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1998-09-24
; PRIOR APPLICATION NUMBER: 60/101,594
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 51
; LENGTH: 428
; TYPE: PRT
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
; OTHER INFORMATION: delta TM
US-09-794-748-51

Query Match          100.0%; Score 16; DB 9; Length 428;
Best Local Similarity 100.0%; Pred. No. 2.6e-07;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 TOHGIRLPLRSGGGA 16
        |||||||
        22 TOHGIRLPLRSGGGA 37

DB      22 TOHGIRLPLRSGGGA 37

RESULT 7
US-09-794-925-51
; Sequence 51, Application US/09794925
; Patent No. US20020064819A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrikson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; TITLE OF INVENTION: THEREFOR
; FILE REFERENCE: 28341/6280HI
; CURRENT FILING DATE: 2001-02-27
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1998-09-24
; PRIOR APPLICATION NUMBER: 60/101,594
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 51
; LENGTH: 428
; TYPE: PRT
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
; OTHER INFORMATION: delta TM
US-09-681-442-51

Query Match          100.0%; Score 16; DB 9; Length 428;
Best Local Similarity 100.0%; Pred. No. 2.6e-07;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 TOHGIRLPLRSGGGA 16
        |||||||
        22 TOHGIRLPLRSGGGA 37

DB      22 TOHGIRLPLRSGGGA 37

RESULT 8
US-09-681-442-51
; Sequence 51, Application US/09681442
; Patent No. US20020081634A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrikson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; TITLE OF INVENTION: THEREFOR
; FILE REFERENCE: 28341/6280FG
; CURRENT FILING DATE: 2001-04-05
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1998-09-24
; PRIOR APPLICATION NUMBER: 60/101,594
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 51
; LENGTH: 428
; TYPE: PRT
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
; OTHER INFORMATION: delta TM
US-09-681-442-51
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Query Match 100.0%; Score 16; DB 9; Length 428;  
Best Local Similarity 100.0%; Pred. No. 2.6e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 TOHGIRLPLRSGLGGA 16  
Db 22 TOHGIRLPLRSGLGGA 37

## RESULT 9

US-09-869-414-51  
; Sequence 51, Application US/09869414  
; Publication No. US20030077226a1  
; GENERAL INFORMATION:  
; APPLICANT: Beinikowski et al.  
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES  
; FILE REFERENCE: 28341/6280M  
; CURRENT FILING DATE: 2001-06-27  
; PRIOR APPLICATION NUMBER: 09/416,901  
; PRIOR FILING DATE: 1999-10-13  
; PRIOR APPLICATION NUMBER: 60/155,493  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: 09/404,133  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: PCT/US99/20881  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: 60/101,594  
; PRIOR FILING DATE: 1998-09-24  
; NUMBER OF SEQ ID NOS: 73  
; SOFTWARE: PatentIn Ver. 2.0  
; SEQ ID NO 51  
; LENGTH: 428  
; TYPE: PRT  
; ORGANISM: Artificial Sequence  
; FEATURE:  
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2(b)  
; OTHER INFORMATION: delta TM  
US-09-869-414-51

Query Match 100.0%; Score 16; DB 10; Length 428;  
Best Local Similarity 100.0%; Pred. No. 2.6e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 TOHGIRLPLRSGLGGA 16  
Db 22 TOHGIRLPLRSGLGGA 37

## RESULT 10

US-09-548-366-51  
; Sequence 51, Application US/09548366  
; Publication No. US20030104365a1  
; GENERAL INFORMATION:  
; APPLICANT: Gurney, Mark E.  
; APPLICANT: Bienkowski, Michael J.  
; APPLICANT: Heinrikson, Robert L.  
; APPLICANT: Parodi, Luis A.  
; APPLICANT: Van, Riqiang  
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND  
; FILE REFERENCE: 28341/6280A  
; CURRENT FILING DATE: 2000-04-12  
; PRIOR APPLICATION NUMBER: 60/155,493  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: 09/404,133  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: PCT/US99/20881  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: 60/101,594  
; PRIOR FILING DATE: 1998-09-24

; NUMBER OF SEQ ID NOS: 65  
; SOFTWARE: PatentIn Ver. 2.0  
; SEQ ID NO 51  
; LENGTH: 428  
; TYPE: PRT  
; ORGANISM: Artificial Sequence  
; FEATURE:  
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2(b)  
; OTHER INFORMATION: delta TM  
US-09-548-366-51

Query Match 100.0%; Score 16; DB 10; Length 428;  
Best Local Similarity 100.0%; Pred. No. 2.6e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 TOHGIRLPLRSGLGGA 16  
Db 22 TOHGIRLPLRSGLGGA 37

## RESULT 11

US-10-652-927-51  
; Sequence 51, Application US/10652927  
; Publication No. US20040043408a1  
; GENERAL INFORMATION:  
; APPLICANT: Gurney et al.  
; TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses  
; FILE REFERENCE: 28915/6280N3  
; CURRENT FILING DATE: 2003-08-29  
; PRIOR APPLICATION NUMBER: 09/794,925  
; PRIOR FILING DATE: 2001-02-27  
; PRIOR APPLICATION NUMBER: 09/416,901  
; PRIOR FILING DATE: 1999-10-13  
; PRIOR APPLICATION NUMBER: 60/155,493  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: 09/404,133  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: PCT/US99/20881  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: 60/101,594  
; PRIOR FILING DATE: 1998-09-24  
; NUMBER OF SEQ ID NOS: 74  
; SOFTWARE: PatentIn Ver. 2.0  
; SEQ ID NO 51  
; LENGTH: 428  
; TYPE: PRT  
; ORGANISM: Artificial sequence  
; FEATURE:  
; OTHER INFORMATION: Hu-Asp2(b) delta TM  
US-10-652-927-51

Query Match 100.0%; Score 16; DB 15; Length 428;  
Best Local Similarity 100.0%; Pred. No. 2.6e-07;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 TOHGIRLPLRSGLGGA 16  
Db 22 TOHGIRLPLRSGLGGA 37

## RESULT 12

US-10-652-830-51  
; Sequence 51, Application US/10652830  
; Publication No. US20040048303a1  
; GENERAL INFORMATION:  
; APPLICANT: Gurney et al.  
; TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses  
; FILE REFERENCE: 28915/6280N1  
; CURRENT FILING DATE: 2003-08-29

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;; PRIOR APPLICATION NUMBER: 09/794,925
;; PRIOR FILING DATE: 2001-02-27
;; PRIOR APPLICATION NUMBER: 09/416,901
;; PRIOR FILING DATE: 1999-10-13
;; PRIOR APPLICATION NUMBER: 60/155,493
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: 09/404,133
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: PCT/US99/20881
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: 60/101,594
;; PRIOR FILING DATE: 1998-09-24
;; NUMBER OF SEQ ID NOS: 74
;; SOFTWARE: Patent In Ver. 2.0
;; SEQ ID NO 51
;; LENGTH: 428
;; TYPE: PRT
;; ORGANISM: Artificial sequence
;; FEATURE:
;; OTHER INFORMATION: Hu-Asp2(b) delta TM
US-10-652-830-51
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Query Match          100.0%; Score 16; DB 15; Length 428;
Best Local Similarity 100.0%; Pred. No. 2.6e-07;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
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```
Oy 1 TOHGIRLPLRSGGGA 16
    |||||
Db 22 TOHGIRLPLRSGGGA 37
```

```
RESULT 13
US-10-652-045-51
;; Sequence 51, Application US/10652045
;; Publication No. US20040166507A1
;; GENERAL INFORMATION:
;; APPLICANT: Gurney et al.
;; TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses
;; FILE REFERENCE: 29915/6280N2
;; CURRENT APPLICATION NUMBER: US/10/652,045
;; CURRENT FILING DATE: 2003-08-29
;; PRIOR APPLICATION NUMBER: 09/794,925
;; PRIOR FILING DATE: 2001-02-27
;; PRIOR APPLICATION NUMBER: 09/416,901
;; PRIOR FILING DATE: 1999-10-13
;; PRIOR APPLICATION NUMBER: 60/155,493
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: 09/404,133
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: PCT/US99/20881
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: 60/101,594
;; PRIOR FILING DATE: 1998-09-24
;; NUMBER OF SEQ ID NOS: 74
;; SOFTWARE: Patent In Ver. 2.0
;; SEQ ID NO 51
;; LENGTH: 428
;; TYPE: PRT
;; ORGANISM: Artificial sequence
;; FEATURE:
;; OTHER INFORMATION: Hu-Asp2(b) delta TM
US-10-652-045-51
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Query Match          100.0%; Score 16; DB 16; Length 428;
Best Local Similarity 100.0%; Pred. No. 2.6e-07;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
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```
Oy 1 TOHGIRLPLRSGGGA 16
    |||||
Db 22 TOHGIRLPLRSGGGA 37
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RESULT 14
US-10-476-935-51
;; Sequence 51, Application US/10476935
;; Publication No. US20040234976A1
;; GENERAL INFORMATION:
;; APPLICANT: Beinkowski et al.
;; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
;; FILE REFERENCE: 28341/6280M1
;; CURRENT APPLICATION NUMBER: US/10/476,935
;; CURRENT FILING DATE: 2003-11-06
;; PRIOR APPLICATION NUMBER: 09/416,901
;; PRIOR FILING DATE: 1999-10-13
;; PRIOR APPLICATION NUMBER: 60/155,493
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: 09/404,133
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: PCT/US99/20881
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: 60/101,594
;; PRIOR FILING DATE: 1998-09-24
;; NUMBER OF SEQ ID NOS: 73
;; SOFTWARE: Patent In Ver. 2.0
;; SEQ ID NO 51
;; LENGTH: 428
;; TYPE: PRT
;; ORGANISM: Artificial Sequence
;; FEATURE:
;; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2(b)
US-10-476-935-51
```

```
Query Match          100.0%; Score 16; DB 16; Length 428;
Best Local Similarity 100.0%; Pred. No. 2.6e-07;
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
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```
Oy 1 TOHGIRLPLRSGGGA 16
    |||||
Db 22 TOHGIRLPLRSGGGA 37
```

```
RESULT 15
US-10-477-076-51
;; Sequence 51, Application US/10477076
;; Publication No. US20050080222A1
;; GENERAL INFORMATION:
;; APPLICANT: Beinkowski et al.
;; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
;; FILE REFERENCE: 28341/6280M2
;; CURRENT APPLICATION NUMBER: US/10/477,076
;; CURRENT FILING DATE: 2003-11-06
;; PRIOR APPLICATION NUMBER: 09/416,901
;; PRIOR FILING DATE: 1999-10-13
;; PRIOR APPLICATION NUMBER: 60/155,493
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: 09/404,133
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: PCT/US99/20881
;; PRIOR FILING DATE: 1999-09-23
;; PRIOR APPLICATION NUMBER: 60/101,594
;; PRIOR FILING DATE: 1998-09-24
;; NUMBER OF SEQ ID NOS: 73
;; SOFTWARE: Patent In Ver. 2.0
;; SEQ ID NO 51
;; LENGTH: 428
;; TYPE: PRT
;; ORGANISM: Artificial Sequence
;; FEATURE:
;; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2(b)
US-10-477-076-51
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Query Match 100.0%; Score 16; DB 17; Length 428;  
 Best Local Similarity 100.0%; Pred. No. 2.6e-07;  
 Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 QY 1 TOHGIRLPLRSGGGA 16  
 |||||  
 Db 22 TOHGIRLPLRSGGGA 37

Search completed: July 26, 2005, 16:41:29  
 Job time : 154 secs

GenCore version 5.1.6  
Copyright (c) 1993 - 2005 CompuGen Ltd.

OM protein - protein search, using sw model

Run on: July 26, 2005, 16:28:04 ; Search time 39 Seconds  
(without alignments)  
39.474 Million cell updates/sec

Title: US-10-726-967A-3

Perfect score: 16

Sequence: 1 TQHGIRLPLRSGLGGA 16

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Gapop 60.0 , Gapext 60.0

Searched: 283416 seqs, 96216763 residues

Word size : 0

Total number of hits satisfying chosen parameters: 283416

Minimum DB seq length: 0

Maximum DB seq length: 200000000

Post-processing: listing first 45 summaries

Database : PIR 79:\*

1: pir1:\*

2: pir2:\*

3: pir3:\*

4: pir4:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Score	Query Match	Length	DB	ID	Description
1	16	100.0	501	2	A59090	aspartic proteinase
2	7	43.8	125	2	E81814	hypothetical prote
3	7	43.8	239	2	AG0420	phosphonates trans
4	7	43.8	252	2	A75547	hypothetical prote
5	7	43.8	352	2	D64966	membrane protein Y
6	6	37.5	157	2	AG2675	hypothetical prote
7	6	37.5	169	2	E90983	probable GDP-L-fuc
8	6	37.5	169	2	H85828	GDP-mannose mannos
9	6	37.5	225	2	D81813	hypothetical prote
10	6	37.5	237	2	T35108	hypothetical prote
11	6	37.5	241	2	T26676	hypothetical prote
12	6	37.5	246	2	AG3644	flagellar biosynth
13	6	37.5	252	2	AE3631	nitrous-oxide redu
14	6	37.5	274	2	D97653	hypothetical prote
15	6	37.5	274	2	AB2877	conserved hypocher
16	6	37.5	284	2	B41224	homeotic protein p
17	6	37.5	292	2	C83520	dihydrodipicolinat
18	6	37.5	299	2	B95149	heat shock protein
19	6	37.5	302	2	A99017	heat shock protein
20	6	37.5	325	2	A72724	hypothetical prote
21	6	37.5	338	2	AE3334	metal chelate tran
22	6	37.5	352	2	T02875	ribosomal protein
23	6	37.5	352	2	D85826	probable transpor
24	6	37.5	352	2	G90980	probable transpor
25	6	37.5	379	2	AF3477	potassium channel
26	6	37.5	390	2	AF3425	oxidoreductase (EC
27	6	37.5	399	2	AC2785	MFS permease (drug
28	6	37.5	399	2	D97564	hypothetical prote
29	6	37.5	411	2	T34585	probable secreted

30	6	37.5	416	2	T32458	hypothetical prote
31	6	37.5	423	2	AA5363	somatoliblerin rece
32	6	37.5	430	2	AB4165	UDP-glucose dehydr
33	6	37.5	430	2	D96536	hypothetical prote
34	6	37.5	437	2	AB0151	conserved hypocher
35	6	37.5	439	2	D70954	hypothetical glyci
36	6	37.5	444	2	B65045	hypothetical prote
37	6	37.5	444	2	C85913	hypothetical prote
38	6	37.5	444	2	A91069	hypothetical prote
39	6	37.5	451	2	I46586	growth hormone-rel
40	6	37.5	454	2	H83377	probable transpor
41	6	37.5	463	2	G83175	probable metallo-o
42	6	37.5	464	2	S29754	growth hormone-rel
43	6	37.5	504	2	AD3629	vdcc protein (limp
44	6	37.5	518	2	S75811	gamma-glutamyltran
45	6	37.5	525	2	AF2950	GDH family prote

#### ALIGNMENTS

RESULT 1  
A59090  
aspartic proteinase (EC 3.4.23.-) BACE precursor - human  
N:Alternate names: beta-secretase; beta-site APP cleaving enzyme  
C:Species: Homo sapiens (man)  
C>Date: 29-Oct-1999 #sequence\_revision 29-Oct-1999 #text\_change 09-Jul-2004  
C/Accession: A59090  
R:Vassar, R.; Bennett, B.D.; Babu-Khan, S.; Kahn, S.; Mendiaz, E.A.; Denis, P.; Teplow, M.A.; Biero, A.L.; Curran, E.; Burgess, T.; Louie, J.C.; Collins, F.; Treanor, J.; Rogers, Science 286, 735-741, 1999  
A>Title: beta-Secretase cleavage of Alzheimer's amyloid precursor protein by the transmembrane  
A:Reference number: A59090; MUID:20002972; PMID:10531052  
A>Note: submitted to Genbank, September 1999  
A/Accession: A59090  
A/Status: not compared with conceptual translation  
A/Molecule type: mRNA  
A:Residues: 1-501 <VAS>  
A:Cross-references: UNIPROT:P56817; GB:AF190725; NID:g6118538; PIDN:AAF04142.1; PID:g611  
C/Genetics:  
A:Gene: BACE  
C:Superfamily: beta-secretase  
C:Keywords: Alzheimer's disease; aspartic proteinase; brain; glycoprotein; hydrolase; pr  
F:1-21/Domain: signal sequence #status predicted <SIG>  
F:22-45/Domain: propeptide #status predicted <PRO>  
F:46-501/Product: acid proteinase BACE #status predicted <MAT>  
F:461-477/Domain: transmembrane #status predicted <TRN>  
F:53,289/Active site: Asp #status predicted  
F:153,172,223,354/Binding site: carbohydrate (Asn) (covalent) #status predicted  
F:330-380/Disulfide bonds: #status predicted

Query Match 100.0%; Score 16; DB 2; Length 501;  
Best Local Similarity 100.0%; Pred. No. 1.8e-09;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 TQHGIRLPLRSGLGGA 16  
DB 22 TQHGIRLPLRSGLGGA 37

RESULT 2  
E81814  
hypothetical protein NMA1874 [imported] - Neisseria meningitidis (strain Z2491 serogrou  
C:Species: Neisseria meningitidis  
C/Date: 05-May-2000 #sequence\_revision 05-May-2000 #text\_change 09-Jul-2004  
C/Accession: E81814  
R:Parkhill, J.; Achtman, M.; James, K.D.; Bentley, S.D.; Churche, C.; Klee, S.R.; Morel  
A:Title: Complete DNA sequence of a serogroup A strain of Neisseria meningitidis Z2491.  
A:Reference number: AB1775; MUID:20222556; PMID:10761919  
A/Accession: E81814  
A/Status: preliminary

A:Molecule type: DNA  
A:Residues: 1-125 <PAR>  
A:Cross-references: UNIPROT:Q9JTC5; GB:AL162757; GB:AL157959; NID:G7380371; PIDN:CAB8509  
A:Experimental source: serogroup A, strain 22491  
C:Genetics:  
A:Gene: NMA1874  
C:Superfamily: Neisseria meningitidis hypothetical protein NMA1874

Query Match 43.8%; Score 7; DB 2; Length 125;  
Best Local Similarity 100.0%; Pred. No. 1.8;  
Matches 7; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 9 LRSGLCG 15  
DB 109 LRSGLCG 115

RESULT 3  
AG0420  
phosphonates transport ATP-binding protein phnL [imported] - Yersinia pestis (strain COG  
C:Species: Yersinia pestis  
C>Date: 02-Nov-2001 #sequence\_revision 02-Nov-2001 #text\_change 09-Jul-2004  
C:Accession: AG0420  
R:Barhill, J.; Wren, B.W.; Thomson, N.R.; Tibball, R.W.; Holden, M.T.G.; Prentice, M.B.  
deno-Farraga, A.M.; Chillingworth, T.; Cronin, A.; Davies, R.M.; Davis, P.; Dougan, G.;  
11, M.; Rutherford, K.; Simmonds, M.; Skelton, J.; Stevens, K.; Whitehead, S.; Barrett,  
Nature 413, 523-527, 2001  
A:Title: Genome sequence of Yersinia pestis, the causative agent of plague.  
A:Reference number: AB0001; MUID:21470413; PMID:11586360  
A:Accession: AG0420  
A:Status: preliminary  
A:Molecule type: DNA  
A:Residues: 1-239 <KUR>  
A:Cross-references: UNIPROT:Q8ZBF4; GB:AL590842; PIDN:CAC92691.1; PID:G15981386; GSPDB:G  
C:Genetics:  
A:Gene: phnL  
C:Superfamily: short-chain ATP-binding cassette proteins; ATP-binding cassette homology

Query Match 43.8%; Score 7; DB 2; Length 239;  
Best Local Similarity 100.0%; Pred. No. 3.2;  
Matches 7; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 2 OHGIRLP 8  
DB 20 OHGIRLP 26

RESULT 4  
A75547  
hypothetical protein - Deinococcus radiodurans (strain R1)  
C:Species: Deinococcus radiodurans  
C>Date: 03-Dec-1999 #sequence\_revision 03-Dec-1999 #text\_change 09-Jul-2004  
C:Accession: A75547  
R:White, O.; Eisen, J.A.; Heidelberg, J.F.; Hickey, E.K.; Peterson, J.D.; Dodson, R.J.;  
S.; Smith, H.O.; Venter, J.C.; Fraser, C.M.  
Science 286, 1571-1577, 1999  
A:Title: Genome sequence of the radioresistant bacterium Deinococcus radiodurans R1.  
A:Reference number: A75540; MUID:20036896; PMID:10567266  
A:Accession: A75547  
A:Status: preliminary  
A:Molecule type: DNA  
A:Residues: 1-259 <WHI>  
A:Cross-references: UNIPROT:Q9PXU2; GB:AE001883; GB:AE000513; NID:G6457878; PIDN:AAF0980  
A:Experimental source: strain R1  
C:Genetics:  
A:Gene: DR0214  
A:Map position: 1  
C:Superfamily: Deinococcus radiodurans hypothetical protein DR0214

Query Match 43.8%; Score 7; DB 2; Length 259;  
Best Local Similarity 100.0%; Pred. No. 3.5;  
Matches 7; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 3 HGIRLPL 9  
DB 190 HGIRLPL 196

RESULT 5  
D64966  
membrane protein yeeB - Escherichia coli (strain K-12)  
C:Species: Escherichia coli  
C>Date: 12-Sep-1997 #sequence\_revision 17-Sep-1997 #text\_change 09-Jul-2004  
C:Accession: D64966  
R:Blattner, F.R.; Plunkett III, G.; Bloch, C.A.; Perna, N.T.; Burland, V.; Riley, M.; Co.  
A.; Rose, D.J.; Mau, B.; Shao, Y.  
Science 277, 1453-1462, 1997  
A:Title: The complete genome sequence of Escherichia coli K-12.  
A:Reference number: A64720; MUID:97426617; PMID:9278503  
A:Accession: D64966  
A:Status: nucleic acid sequence not shown; translation not shown  
A:Molecule type: DNA  
A:Residues: 1-352 <BIAT>  
A:Cross-references: UNIPROT:P33015; GB:AE000292; GB:U00096; NID:G1788310; PIDN:AACT5074.1  
A:Experimental source: strain K-12, substrain MG1655  
C:Genetics:  
A:Gene: yeeB  
C:Keywords: transmembrane protein  
F:2-18/Domain: transmembrane #status predicted <TM1>  
F:44-60/Domain: transmembrane #status predicted <TM2>  
F:75-91/Domain: transmembrane #status predicted <TM3>  
F:106-122/Domain: transmembrane #status predicted <TM4>  
F:153-169/Domain: transmembrane #status predicted <TM5>  
F:201-217/Domain: transmembrane #status predicted <TM6>  
F:250-266/Domain: transmembrane #status predicted <TM7>  
F:320-336/Domain: transmembrane #status predicted <TM8>

Query Match 43.8%; Score 7; DB 2; Length 352;  
Best Local Similarity 100.0%; Pred. No. 4.6;  
Matches 7; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 9 LRSGLCG 15  
DB 283 LRSGLCG 289

RESULT 6  
AG2675  
hypothetical protein Atu0805 [imported] - Agrobacterium tumefaciens (strain C58, Dupont)  
C:Species: Agrobacterium tumefaciens  
C>Date: 11-Jan-2002 #sequence\_revision 11-Jan-2002 #text\_change 09-Jul-2004  
C:Accession: AG2675  
R:Wood, D.W.; Setubal, J.C.; Kaul, R.; Monks, D.; Chen, L.; Wood, G.E.; Chen, Y.; Woo, L.  
erge, G.; Gilliet, W.; Grant, C.; Guenther, D.; Kutayavin, T.; Levy, R.; Li, M.; McTiell,  
Science 294, 2317-2323, 2001  
A:Authors: Yoo, H.; Tao, Y.; Biddle, P.; Jung, M.; Kreepan, W.; Perry, M.; Gordon-Kamm, I.  
ster, E.W.  
A:Title: The Genome of the Natural Genetic Engineer Agrobacterium tumefaciens C58.  
A:Reference number: AB2577; MUID:21608550; PMID:11743193  
A:Accession: AG2675  
A:Status: preliminary  
A:Molecule type: DNA  
A:Residues: 1-157 <KUR>  
A:Cross-references: UNIPROT:O8UH78; GB:AE006688; PIDN:AL41821.1; PID:G17739178; GSPDB:G  
A:Experimental source: strain C58 (Dupont)  
C:Genetics:  
A:Gene: Atu0805  
A:Map position: circular chromosome

Query Match 37.5%; Score 6; DB 2; Length 157;  
Best Local Similarity 100.0%; Pred. No. 25;  
Matches 6; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 8 PLRSGL 13



Db 81 PLRSGL 86

## RESULT 7

E90983

probable GDP-L-fucose pathway enzyme [imported] - Escherichia coli (strain O157:H7, sub  
C:Species: Escherichia coli  
C:Date: 18-Jul-2001 #sequence\_revision 18-Jul-2001 #text\_change 16-Aug-2004  
C:Accession: E90983

R:Hayashi, T.; Makino, K.; Onishi, M.; Kurokawa, K.; Ishii, K.; Yokoyama, K.; Han, C.G.  
gasawara, N.; Yasunaga, T.; Kuhara, S.; Shiba, T.; Hattori, M.; Shinagawa, H.

DNA Res. 8, 11-22, 2001

A:Title: Complete genome sequence of enterohemorrhagic Escherichia coli O157:H7 and genc  
A:Reference number: A99629; MUID:21156231; PMID:11258796

A:Accession: E90983

A:Status: preliminary

A:Molecule type: DNA

A:Residues: 1-169 &lt;HAV&gt;

A:Cross-references: UNIPROT:O85341; GB:BA000007; PIDN:BA036260.1; PID:g13362305; GSPDB:G  
A:Experimental source: strain O157:H7, substrain RMD 0509952

C:Genetics:

A:Gene: EC62837

C:Superfamily: mut domain homology

Query Match 37.5%; Score 6; DB 2; Length 169;  
Best Local Similarity 100.0%; Pred. No. 27;

Matches 6; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 4 GIRLPL 9

Db 71 GIRLPL 76

## RESULT 8

H85828

GDP-mannose mannosylhydrolase [imported] - Escherichia coli (strain O157:H7, substrain H  
C:Species: Escherichia coli  
C:Date: 16-Feb-2001 #sequence\_revision 16-Feb-2001 #text\_change 16-Aug-2004  
C:Accession: H85828

R:Perna, N.T.; Plunkett III, G.; Burland, V.; Mau, B.; Glasner, J.D.; Rose, D.J.; Mayhew

iller, L.; Grobeck, E.J.; Davis, N.W.; Lam, A.; Dimalanta, E.; Potamoudis, K.; Apodaca,  
Nature 409, 529-533, 2001

A:Title: Genome sequence of enterohemorrhagic Escherichia coli O157:H7.

A:Reference number: AB5480; MUID:21074935; PMID:11206551

A:Accession: H85828

A:Status: preliminary

A:Molecule type: DNA

A:Residues: 1-169 &lt;STO&gt;

A:Cross-references: UNIPROT:O85341; GB:AE005174; NID:g12516220; PIDN:AA057092.1; GSPDB:G  
A:Experimental source: strain O157:H7, substrain EDL933

C:Genetics:

A:Gene: wbdQ

C:Superfamily: mut domain homology

Query Match 37.5%; Score 6; DB 2; Length 169;  
Best Local Similarity 100.0%; Pred. No. 27;

Matches 6; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 4 GIRLPL 9

Db 71 GIRLPL 76

## RESULT 9

D81813

hypothetical protein NMA1865 [imported] - Neisseria meningitidis (strain Z2491 serogroup  
C:Species: Neisseria meningitidis  
C:Date: 05-May-2000 #sequence\_revision 05-May-2000 #text\_change 09-Jul-2004  
C:Accession: D81813

R:Parkhill, J.; Achtman, M.; James, K.D.; Bentley, S.D.; Churcher, C.; Klee, S.R.; Morel  
; Holroyd, S.; Jagers, K.; Leather, S.; Moule, S.; Mungall, K.; Quail, M.A.; Rajandream,  
Nature 404, 502-506, 2000

A:Title: Complete DNA sequence of a serogroup A strain of Neisseria meningitidis Z2491.

A:Reference number: A81775; MUID:20222556; PMID:10761919

A:Accession: D81813

A:Status: preliminary

A:Molecule type: DNA

A:Residues: 1-225 &lt;PAR&gt;

A:Cross-references: UNIPROT:O9J968; GB:AL162757; GB:AL157959; NID:g7380371; PIDN:CAB8508

A:Experimental source: serogroup A, strain Z2491

C:Genetics:

A:Gene: NMA1865

Query Match 37.5%; Score 6; DB 2; Length 225;  
Best Local Similarity 100.0%; Pred. No. 35;

Matches 6; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 4 GIRLPL 9

Db 197 GIRLPL 202

## RESULT 10

T35108

hypothetical protein SC4H2.09 SC4H2.09 - Streptomyces coelicolor  
C:Species: Streptomyces coelicolor  
C:Date: 05-Nov-1999 #sequence\_revision 05-Nov-1999 #text\_change 09-Jul-2004  
C:Accession: T35108

R:Seeger, K.J.; Harris, D.; Parkhill, J.; Barrell, B.G.; Rajandream, M.A.  
submitted to the EMBL Data Library, March 1998

A:Reference number: Z21568

A:Accession: T35108

A:Status: preliminary; translated from GB/EMBL/DBJ

A:Molecule type: DNA

A:Residues: 1-237 &lt;SEE&gt;

A:Cross-references: UNIPROT:O69964; EMBL:AL022268; PIDN:CAA18325.1; GSPDB:GN00070; SCOD

A:Experimental source: strain A3(2)

C:Genetics:

A:Gene: SCODP:SC4H2.09

C:Superfamily: Streptomyces coelicolor hypothetical protein SC4H2.09

Query Match 37.5%; Score 6; DB 2; Length 237;  
Best Local Similarity 100.0%; Pred. No. 37;

Matches 6; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 11 SGLGGA 16

Db 214 SGLGGA 219

## RESULT 11

T26676

hypothetical protein Y38F1A.1 - Caenorhabditis elegans  
C:Species: Caenorhabditis elegans  
C:Date: 15-Oct-1999 #sequence\_revision 15-Oct-1999 #text\_change 09-Jul-2004  
C:Accession: T26676

R:Wallis, J.

submitted to the EMBL Data Library, October 1998

A:Reference number: Z20253

A:Accession: T26676

A:Status: preliminary; translated from GB/EMBL/DBJ

A:Molecule type: DNA

A:Residues: 1-241 &lt;WIL&gt;

A:Cross-references: UNIPROT:O9XWMS; EMBL:AL032639; PIDN:CAA21628.1; GSPDB:GN00020; CESP:

A:Experimental source: clone Y38F1A

C:Genetics:

A:Gene: CESP:Y38F1A.1

A:Map position: 2

A:Introns: 37/3; 76/2; 130/3; 202/3

Query Match 37.5%; Score 6; DB 2; Length 241;  
Best Local Similarity 100.0%; Pred. No. 37;

Matches 6; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 11 SGLGGA 16

Db 101 SGLGGA 106

# RESULT 12

AG3644

flagellar biosynthetic protein flp [imported] - Brucella melitensis (strain 16M)

C/Species: Brucella melitensis

C/Date: 01-Feb-2002 #sequence\_revision 01-Feb-2002 #text\_change 09-Jul-2004

C/Accession: AG3644

R/DeVecchio, V.G.; Kapral, V.; Redkar, R.U.; Patra, G.; Mujer, C.; Ios, T.; Ivanova,

; Mazur, M.; Goldsman, E.; Selkov, E.; Elizer, P.H.; Hagius, S.; O'Callaghan, D.; Leles

Proc. Natl. Acad. Sci. U.S.A. 99, 443-448, 2002

A/Title: The genome sequence of the facultative intracellular pathogen Brucella melit

A/Reference number: AD3252; PMID:11756688

A/Accession: AG3644

A/Status: preliminary

A/Molecule type: DNA

A/Residues: 1-246 <KUR>

A/Cross-references: UNIPROT:Q8YB21; GB:AE008918; PIDN:AAL54322.1; PID:gl7985302; GSPDB:C

A/Experimental source: strain 16M

C/Genetics:

A/Gene: BME11080

A/Map position: 11

C/Superfamily: flagellar biosynthetic protein flp

Query Match 37.5%; Score 6; DB 2; Length 246;

Best Local Similarity 100.0%; Pred. No. 38;

Matches 6; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 9 RSGG 14

Db 75 RSGG 80

# RESULT 13

AE3631

nitrous-oxide reductase (EC 1.7.99.6) [imported] - Brucella melitensis (strain 16M)

C/Species: Brucella melitensis

C/Date: 01-Feb-2002 #sequence\_revision 01-Feb-2002 #text\_change 09-Jul-2004

C/Accession: AE3631

R/DeVecchio, V.G.; Kapral, V.; Redkar, R.U.; Patra, G.; Mujer, C.; Ios, T.; Ivanova,

; Mazur, M.; Goldsman, E.; Selkov, E.; Elizer, P.H.; Hagius, S.; O'Callaghan, D.; Leles

Proc. Natl. Acad. Sci. U.S.A. 99, 443-448, 2002

A/Title: The genome sequence of the facultative intracellular pathogen Brucella melit

A/Reference number: AD3252; PMID:11756688

A/Accession: AE3631

A/Status: preliminary

A/Molecule type: DNA

A/Residues: 1-252 <KUR>

A/Cross-references: UNIPROT:Q8YB21; GB:AE008918; PIDN:AAL54216.1; PID:gl7985186; GSPDB:C

A/Experimental source: strain 16M

C/Genetics:

A/Gene: BME10974

A/Map position: 11

C/Keywords: oxidoreductase

Query Match 37.5%; Score 6; DB 2; Length 252;

Best Local Similarity 100.0%; Pred. No. 39;

Matches 6; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 11 SGLGGA 16

Db 28 SGLGGA 33

# RESULT 14

D97653

hypothetical protein AGR\_C\_4436 [imported] - Agrobacterium tumefaciens (strain C58, Cere

C/Species: Agrobacterium tumefaciens

C/Date: 30-Sep-2001 #sequence\_revision 30-Sep-2001 #text\_change 09-Jul-2004

C/Accession: D97653

R/Goodner, B.; Hinkle, G.; Gattung, S.; Miller, N.; Blanchard, M.; Qurollo, B.; Goldman,

A.; Liu, F.; Wollam, C.; Allinger, M.; Doughy, D.; Scott, C.; Lappas, C.; Markelz, B.; Science 294, 2323-2328, 2001

A/Title: Genome Sequence of the Plant Pathogen and Biotechnology Agent Agrobacterium tume

A/Reference number: A97359; MUID:21608551; PMID:11743194

A/Accession: D97653

A/Status: preliminary

A/Molecule type: DNA

A/Residues: 1-274 <KUR>

A/Cross-references: UNIPROT:Q8UCN9; GB:AE007869; PIDN:AAK8181.1; PID:gl5157627; GSPDB:G

C/Genetics:

A/Gene: AGR\_C\_4436

A/Map position: circular chromosome

Query Match 37.5%; Score 6; DB 2; Length 274;

Best Local Similarity 100.0%; Pred. No. 42;

Matches 6; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 10 RSGG 15

Db 249 RSGG 254

# RESULT 15

AB2877

conserved hypothetical protein Atu2444 [imported] - Agrobacterium tumefaciens (strain C58)

C/Species: Agrobacterium tumefaciens

C/Date: 11-Jan-2002 #sequence\_revision 11-Jan-2002 #text\_change 09-Jul-2004

C/Accession: AB2877

R/Wood, D.W.; Setubal, J.C.; Kaul, R.; Monke, D.; Chen, L.; Wood, G.E.; Chen, Y.; Woo, L

erage, G.; Gillet, W.; Grant, C.; Guenther, D.; Kutayvin, T.; Levy, R.; Li, M.; McClellan

; Karp, P.; Romero, P.; Zhang, S. Science 294, 2317-2323, 2001

A/Authors: Yoo, H.; Tao, Y.; Biddle, P.; Jung, M.; Krespan, W.; Perry, M.; Gordon-Kamm,

ster, E.W.

A/Title: The Genome of the Natural Genetic Engineer Agrobacterium tumefaciens C58.

A/Reference number: AB2577; MUID:21608550; PMID:11743193

A/Accession: AB2877

A/Status: preliminary

A/Molecule type: DNA

A/Residues: 1-274 <KUR>

A/Cross-references: UNIPROT:Q8UCN9; GB:AE008688; PIDN:AAL43432.1; PID:gl7740934; GSPDB:C

A/Experimental source: strain C58 (Dupont)

C/Genetics:

A/Gene: Atu2444

A/Map position: circular chromosome

Query Match 37.5%; Score 6; DB 2; Length 274;

Best Local Similarity 100.0%; Pred. No. 42;

Matches 6; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 10 RSGG 15

Db 249 RSGG 254

Search completed: July 26, 2005, 16:38:03  
Job time : 41 secs

GenCore version 5.1.6  
Copyright (c) 1993 - 2005 CompuGen Ltd.

OM protein - protein search, using sw model

Run on: July 26, 2005, 16:27:19 ; Search time 173 Seconds  
(without alignments)  
47.360 Million cell updates/sec

Title: US-10-726-967A-3  
Perfect score: 16  
Sequence: 1 TQHGIRLPRLRSGLGCA 16

Scoring table: OLIGO  
Gapop 60.0 , Gapept 60.0

Searched: 1612378 seqs, 512079187 residues

Word size : 0

Total number of hits satisfying chosen parameters: 1612378

Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Listing first 45 summaries

Database : Uniprot 03: \*  
1: uniprot\_sprot: \*  
2: uniprot\_trembl: \*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	16	100.0	127	2	Q76KP0
2	16	100.0	501	1	BAEI_HUMAN
3	16	100.0	501	2	Q81YC8
4	10	62.5	467	2	Q8C4P4
5	10	62.5	501	1	BAEI_MOUSE
6	10	62.5	501	1	BAEI_RAT
7	10	62.5	501	2	Q8B0V4
8	10	62.5	501	2	Q8C7R1
9	8	50.0	814	2	Q7S2I8
10	7	43.8	97	2	Q9JPF6
11	7	43.8	125	2	Q9UTC5
12	7	43.8	181	2	Q7NTR4
13	7	43.8	239	2	Q6E2F9
14	7	43.8	239	2	Q8ZBF4
15	7	43.8	259	2	Q9RXU2
16	7	43.8	272	2	Q67QZ3
17	7	43.8	293	2	Q74X41
18	7	43.8	299	2	Q9JHS7
19	7	43.8	304	2	Q7SFA9
20	7	43.8	329	2	Q7NTO7
21	7	43.8	350	2	Q9JF54
22	7	43.8	352	1	YEEB_ECOLI
23	7	43.8	352	2	Q9JF48
24	7	43.8	352	2	Q9JF55
25	7	43.8	352	2	Q9JF57
26	7	43.8	352	2	Q9JF60
27	7	43.8	352	2	Q83R10
28	7	43.8	353	2	Q9JF51
29	7	43.8	354	2	Q9JF52
30	7	43.8	354	2	Q9JF61
31	7	43.8	457	2	Q8B6W9

32	7	43.8	460	2	Q9X3V2	Q9X3V2 pseudomonas
33	7	43.8	468	2	Q8BPI9	Q8BPI9 pseudomonas
34	7	43.8	629	2	Q67QI5	Q67QI5 symbiodace
35	7	43.8	889	2	Q42723	Q42723 emericella
36	7	43.8	911	1	CAFA_MOUSE	Q9GWF0 mus musculu
37	7	43.8	1104	2	Q9F060	Q9F060 oryza sativ
38	6	37.5	55	2	Q8PKC1	Q8PKC1 xanthomonas
39	6	37.5	55	2	Q7UVPI	Q7UVPI rhodospirill
40	6	37.5	61	2	Q7S3N8	Q7S3N8 neurospora
41	6	37.5	68	2	Q6J3X9	Q6J3X9 burkholderi
42	6	37.5	77	2	Q7AET6	Q7AET6 geobacter s
43	6	37.5	99	2	Q949C5	Q949C5 oryza sativ
44	6	37.5	99	2	Q6H7G5	Q6H7G5 oryza sativ
45	6	37.5	100	2	Q6ZB08	Q6ZB08 oryza sativ

## ALIGNMENTS

RESULT 1  
ID Q76KP0 PRELIMINARY; PRT; 127 AA.  
AC Q76KP0;  
DT 05-JUL-2004 (TREMBLrel. 27, Created)  
DT 05-JUL-2004 (TREMBLrel. 27, Last sequence update)  
DT 05-JUL-2004 (TREMBLrel. 27, Last annotation update)  
DE Beta-site APP cleaving enzyme isoform I-127.  
GN Name=BACE;  
OS Homo sapiens (Human).  
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
OC Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.  
OX NCBI\_Taxid=9606;  
RN [1]  
RP SEQUENCE FROM N.A.  
RA Tanahashi H.;  
RL Submitted (Aug-2002) to the EMBL/GenBank/DBJ databases.  
CC -1 SIMILARITY: Belongs to the EMBL/GenBank/DBJ databases.  
DR EMBL; AB089958; BAC81826.1; -  
DR HSSP; P00797; 1BBS.  
DR GO; GO:0009049; F:aspartic-type signal peptidase activity; IEA.  
DR GO; GO:000508; P:proteolysis and peptidolysis; IEA.  
DR InterPro; IPR001461; Peptidase\_A1.  
DR InterPro; IPR009119; Pept\_A1\_BACE.  
DR InterPro; IPR009120; Pept\_A1\_BACE.  
DR InterPro; IPR009007; Pept\_Aspartic.  
DR InterPro; IPR001969; Pept\_Asp\_AS.  
DR Pfam; PF00026; Asp\_1.  
DR PRINTS; PR01816; BACE1.  
DR PRINTS; PR01815; BACEFAMILY.  
DR PROSITE; PS00141; ASP\_PROTEASE; 1.  
KW Aspartyl protease; Hydrolase; Protease.  
SQ SEQUENCE 127 AA; 1393 MW; C657354CBEE72DC4 CRC64;  
Query Match 100.0%; Score 16; DB 2; Length 127;  
Best Local Similarity 100.0%; Pred. No. 1.3e-08;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
QY 1 TQHGIRLPRLRSGLGCA 16  
Db 22 TQHGIRLPRLRSGLGCA 37  
RESULT 2  
ID BAEI\_HUMAN STANDARD; PRT; 501 AA.  
AC P56817; Q9BYB9; Q9BYC0; Q9BYC1; Q9JUT5;  
DT 30-MAY-2000 (Rel. 39, Created)  
DT 30-MAY-2000 (Rel. 39, Last sequence update)  
DT 25-OCT-2004 (Rel. 45, Last annotation update)  
DE Beta-secretase 1 precursor (EC 3.4.23.46) (Beta-site APP cleaving enzyme 1) (Beta-site amyloid precursor protein cleaving enzyme 1) (Aspartyl protease 2) (Aap 2) (ASP2) (Membrane-associated aspartic protease 2) (Memapsin-2).  
DE

GN Name=BACE1; Synonyms=BACE;  
 OS Homo sapiens (Human).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 CC Mammalia; Eutheria; Primates; Catarrhini; Hominiidae; Homo.  
 CX NCBI\_TaxID=9606;  
 RN [1]  
 RP SEQUENCE FROM N.A. (ISOFORM A).  
 RC TISSUE=Brain;  
 RX MEDLINE=20002972; PubMed=10531052; DOI=10.1126/science.286.5440.735;  
 RA Vassar R., Bennett B.D., Babu-Khan S., Kahn S., Mendiaz E.A.,  
 RA Denis P., Teplow D.B., Ross S., Amaratne P., Loebner R., Luo Y.,  
 RA Fisher P., Fuller J., Edenson S., Lile J., Jarosinski M.A.,  
 RA Biere A.U., Curran E., Burgess T., Louis J.-C., Collins F.,  
 RA Treanor J., Rogers G., Citron M.;  
 RT "Beta-secretase cleavage of Alzheimer's amyloid precursor protein by  
 RT the transmembrane aspartic protease BACE.";  
 RL Science 286:735-741(1999).  
 RN [2]  
 RP SEQUENCE FROM N.A. (ISOFORM A), SEQUENCE OF 46-68, AND  
 RC CHARACTERIZATION.  
 RX TISSUE=Brain;  
 RX MEDLINE=20057171; PubMed=10591214; DOI=10.1038/990114;  
 RA Sinha S., Anderson J.P., Barbour R., Basl G.S., Caccavello R.,  
 RA Davis D., Dean M., Dovey H.F., Frigon N., Hong J., Jacobson-Croak K.,  
 RA Jewett N., Keim P., Knops J., Lieberburg I., Power M., Tan H.,  
 RA Tatsuno G., Tung J., Schenk D., Seubert P., Suenensaat S.M., Wang S.,  
 RA Walker D., Zhao J., McConlogue L., Varghese J.;  
 RT "Purification and cloning of amyloid precursor protein beta-secretase  
 RT from human brain.";  
 RL Nature 402:537-540(1999).  
 RN [3]  
 RP SEQUENCE FROM N.A. (ISOFORM A).  
 RX MEDLINE=20057170; PubMed=10591213; DOI=10.1038/990107;  
 RA Yan R., Benkowitz M.J., Shuck M.E., Miao H., Tory M.C., Pauley A.M.,  
 RA Yan R., Benkowitz M.J., Shuck M.E., Miao H., Tory M.C., Pauley A.M.,  
 RA Tomashvili A.G., Parodi L.A., Heinrichson R.L., Gurney M.B.;  
 RT "Membrane-anchored aspartyl protease with Alzheimer's disease beta-  
 RT secretase activity.";  
 RL Nature 402:533-537(1999).  
 RN [4]  
 RP SEQUENCE FROM N.A. (ISOFORM A).  
 RX MEDLINE=20120043; PubMed=10656250; DOI=10.1006/mcne.1999.0811;  
 RA Hussain I., Powell D.J., Howlett D.R., Tew D.G., Meek T.D.,  
 RA Chapman C., Gloger I.S., Murphy K.E., Southern C.D., Ryan D.M.,  
 RA Smith T.S., Simmons D.L., Walsh F.S., Dingwall C., Christie G.;  
 RT "Identification of a novel aspartic protease (Asp 2) as beta-  
 RT secretase.";  
 RL Mol. Cell. Neurosci. 14:419-427(1999).  
 RN [5]  
 RP SEQUENCE FROM N.A. (ISOFORM B).  
 RC TISSUE=Brain, and Pancreas;  
 RA Michel B., De Pietri Tonelli D., Zaccchetti D., Keller P.;  
 RT "New beta-site APP cleaving enzyme isoform (BACE-1B) obtained from  
 RT human brain and pancreas.";  
 RL Submitted (JAN-2001) to the EMBL/GenBank/DBJ databases.  
 RN [6]  
 RP SEQUENCE FROM N.A. (ISOFORM C).  
 RC TISSUE=Pancreas;  
 RA Zaccchetti D., De Pietri Tonelli D., Schnurbus R.;  
 RT "New beta-site APP cleaving enzyme isoform (BACE-1C) obtained from  
 RT human pancreas.";  
 RL Submitted (JAN-2001) to the EMBL/GenBank/DBJ databases.  
 RN [7]  
 RP SEQUENCE FROM N.A. (ISOFORMS B, C AND D).  
 RC TISSUE=Brain;  
 RX MEDLINE=21408467; PubMed=11516566; DOI=10.1016/S0304-3940(01)01912-7;  
 RA Tanahashi H., Tsubita T.;  
 RT "Three novel alternatively spliced isoforms of the human beta-site  
 RT amyloid precursor protein cleaving enzyme (BACE) and their effect on  
 RT Neurosci. Lett. 307:9-12(2001).  
 RN [8]  
 RP SEQUENCE OF 14-501 FROM N.A. (ISOFORM A), AND CHARACTERIZATION.

RX MEDLINE=20144060; PubMed=10677483; DOI=10.1073/pnas.97.4.1456;  
 RA Lin X., Koelsch G., Wu S., Downs D., Dabhi A., Tang J.;  
 RT "Human aspartic protease memapsin 2 cleaves the beta-secretase site of  
 RT beta-amyloid precursor protein.";  
 RL Proc. Natl. Acad. Sci. U.S.A. 97:1456-1460(2000).  
 RN [9]  
 RP DISULFIDE BONDS.  
 RX MEDLINE=21950860; PubMed=11953458;  
 RA Fischer F., Molinari M., Bodendorf U., Paganetti P.;  
 RT "The disulphide bonds in the catalytic domain of BACE are critical but  
 RT not essential for amyloid precursor protein processing activity.";  
 RL Neurochem. 80:1079-1086(2002).  
 CC -1- FUNCTION: Responsible for the proteolytic processing of the  
 CC amyloid precursor protein (APP). Cleaves at the amino terminus of  
 CC the A-beta peptide sequence, between residues 671 and 672 of APP,  
 CC leads to the generation and extracellular release of beta-cleaved  
 CC soluble APP, and a corresponding cell-associated carboxy-terminal  
 CC fragment which is later released by gamma-secretase.  
 CC -1- CATALYTIC ACTIVITY: Broad endopeptidase specificity. Cleaves Glu-  
 CC Val-Ileu-Ile-Asp-Ala-Glu-Phe in the Swedish variant of  
 CC Alzheimer's amyloid precursor protein.  
 CC -1- SUBCELLULAR LOCATION: Type I membrane protein.  
 CC -1- ALTERNATIVE PRODUCTS:  
 CC Event=Alternative splicing; Named isoforms=4;  
 CC Name=A; Synonyms=BACE-1A, BAC-501,  
 CC IsoId=P56817-1; Sequence=D1sp1ayd;  
 CC Name=B; Synonyms=BACE-1B, BACE-1-476;  
 CC IsoId=P56817-2; Sequence=VSP\_005223;  
 CC Name=C; Synonyms=BACE-1C, BACE-1-457;  
 CC IsoId=P56817-3; Sequence=VSP\_005222;  
 CC Name=D; Synonyms=BACE-1D, BACE-1-432;  
 CC IsoId=P56817-4; Sequence=VSP\_005222; VSP\_005223;  
 CC -1- TISSUE SPECIFICITY: Brain.  
 CC -1- SIMILARITY: Belongs to the peptidase A1 family.  
 CC -----  
 CC This SWISS-PROT entry is copyright. It is produced through a collaboration  
 CC between the Swiss Institute of Bioinformatics and the EMBL collaboration  
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 CC or send an email to [license@isb-sib.ch](mailto:license@isb-sib.ch)).  
 CC -----  
 DR EMBL; AF190725; AAF04142.1; -  
 DR EMBL; AF201468; AAF18982.1; -  
 DR EMBL; AF200343; AAF17079.1; -  
 DR EMBL; AF204943; AAF26367.1; -  
 DR EMBL; AF338816; AAK38374.1; -  
 DR EMBL; AF338817; AAK38375.1; -  
 DR EMBL; AB050436; BAB40931.1; -  
 DR EMBL; AB050437; BAB40932.1; -  
 DR EMBL; AB050438; BAB40933.1; -  
 DR EMBL; AF200193; AAF13715.1; -  
 DR PIR; A59090; A59090;  
 DR PDB; 1FKN; X-ray; A/B=56-446.  
 DR PDB; 1M4H; X-ray; A/B=56-446.  
 DR MEROPS; A01.004; -  
 DR GeneW; HGNC:933; BACE1.  
 DR H-InvDB; HIX0010165; -  
 DR MIM; 604252; -  
 DR GO; GO:0005887; C:integral to plasma membrane; TAS.  
 DR GO; GO:0008796; F:beta-aspartyl-peptidase activity; TAS.  
 DR GO; GO:0006508; P:proteolysis and peptidolysis; TAS.  
 DR InterPro; IPR009119; Pept\_A1\_BACE.  
 DR InterPro; IPR009120; Pept\_A1\_BACE.  
 DR InterPro; IPR001969; Pept\_Asp\_AS.  
 DR InterPro; IPR009007; Pept\_AspArtic.  
 DR InterPro; IPR001461; Peptidase\_A1.  
 DR Pfam; PF00026; Asp\_1.  
 DR PRINTS; PRO1816; BACE1.  
 DR PRINTS; PRO1815; BACEFAMILY.  
 DR PRINTS; PRO0792; PEPsin.  
 DR PROSITE; PS00141; ASP\_PROTEASE; 1.

KW 3D-structure; Alternative splicing; Aspartyl protease;  
KW Direct protein sequencing; Glycoprotein; Hydrolase; Signal;  
KW Transmembrane; Zymogen.  
FT SIGNAL 1 21 Potential.  
FT PROPEP 22 45 Beta-secretase 1.  
FT CHAIN 46 501 Extracellular (Potential).  
FT DOMAIN 22 457 Potential.  
FT TRANSMEM 458 478 Cytoplasmic (Potential).  
FT DOMAIN 479 501 By similarity.  
FT ACT SITE 93 93 By similarity.  
FT ACT SITE 289 289 By similarity.  
FT DISULFID 216 420  
FT DISULFID 278 443  
FT DISULFID 330 380  
FT CARBOHYD 153 153 N-linked (GlcNAc... ) (Potential).  
FT CARBOHYD 172 172 N-linked (GlcNAc... ) (Potential).  
FT CARBOHYD 223 223 N-linked (GlcNAc... ) (Potential).  
FT CARBOHYD 354 354 N-linked (GlcNAc... ) (Potential).  
FT VARSPLIC 146 189 Missing (in isoform C and isoform D).  
FT VARSPLIC 190 214 /FTid=VSP\_005222.  
FT VARSPLIC 214 Missing (in isoform B and isoform D).  
FT VARSPLIC 214 /FTid=VSP\_005223.  
FT HELIX 61 63  
FT TURN 64 65  
FT STRAND 67 70  
FT TURN 71 73  
FT STRAND 74 81  
FT TURN 82 85  
FT STRAND 86 93  
FT TURN 94 95  
FT STRAND 99 102  
FT TURN 107 108  
FT HELIX 115 117  
FT TURN 119 120  
FT STRAND 122 131  
FT STRAND 136 147  
FT TURN 149 150  
FT STRAND 155 167  
FT TURN 172 173  
FT STRAND 178 181  
FT HELIX 185 187  
FT TURN 192 193  
FT HELIX 197 204  
FT STRAND 211 215  
FT HELIX 224 229

Query Match 100.0%; Score 16; DB 1; Length 501;  
Best Local Similarity 100.0%; Pred. No. 4.2e-08;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 TOHGIRLPLRSGIGGA 16  
DB 22 TOHGIRLPLRSGIGGA 37

RESULT 3  
Q8IYC8 PRELIMINARY; PRT; 501 AA.  
ID Q8IYC8  
AC Q8IYC8  
DT 01-MAR-2003 (TrEMBLrel. 23, Created)  
DT 01-MAR-2003 (TrEMBLrel. 23, Last sequence update)  
DT 01-MAR-2004 (TrEMBLrel. 26, Last annotation update)  
DE Beta-site APP-cleaving enzyme 1, isoform A preproprotein.  
GN Name=BACE1;  
OS Homo sapiens (Human).  
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
OC Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.  
OC NCB1\_TaxID=9606;  
OX NCB1  
RN [1]  
RP SEQUENCE FROM N.A.  
RC TISSUE=Brain;  
RX MEDLINE=22388257; PubMed=12477932; DOI=10.1073/pnas.242603899;  
RA Strausberg R.L., Feingold E.A., Grouse L.H., Derge J.G.,

RA Klausner R.D., Collins F.S., Wagner L., Sherman C.M., Schuler G.D.,  
RA Altschul S.F., Zeeberg B., Buetow K.H., Schaefer C.F., Bhat N.K.,  
RA Hopkins R.F., Jordan H., Moore T., Max S.I., Wang J., Hsieh F.,  
RA Diatchenko L., Marusina K., Farmer A.A., Rubin G.M., Hong L.,  
RA Stapleton M., Soares M.B., Bonaldo M.F., Casavant T.L., Scheetz T.E.,  
RA Brownstein M.J., Usdin T.B., Toshiyuki S., Carrinci P., Prange C.,  
RA Raha S.S., Loguella N.A., Peters G.J., Abramson R.D., Mullany S.J.,  
RA Bosak S.A., McGowan P.J., McKernan K.J., Malek J.A., Gunaratne P.H.,  
RA Richards S., Morley K.C., Hale S., Garcia A.M., Gay L.J., Huiyk S.W.,  
RA Villalón D.K., Muzny D.M., Sodergren E.J., Lu X., Gibbs R.A.,  
RA Fanev J., Helton E., Kerteman M., Madan A., Rodriguez T.L., Sanchez A.,  
RA Whiting M., Madan A., Young A.C., Shevchenko Y., Bouffard G.G.,  
RA Blakesley R.W., Touchman J.W., Green E.D., Dickson M.C.,  
RA Rodriguez A.C., Grimwood J., Schmutz J., Myers R.M., Butlerfield Y.S.,  
RA Krzywinski M.I., Skalska U., Smailus D.E., Scherch A., Schein J.E.,  
RA Jones S.J., Marra M.A.;  
RT "Generation and initial analysis of more than 15,000 full-length human  
RT and mouse cDNA sequences";  
RL Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903 (2002).  
RN [2]  
RP SEQUENCE FROM N.A.  
RC TISSUE=Brain;  
RA Strausberg R.;  
RL Submitted (JUL-2002) to the EMBL/Genbank/DBJ databases.  
CC -1 SIMILARITY: Belongs to peptidase family A1.  
DR EMBL; BC036084; AAH36084.1; -.  
DR HSP; P56817; 1PKN.  
DR GO; GO:0005768; Cytoplasm; ISS.  
DR GO; GO:0005794; C:Golgi apparatus; ISS.  
DR GO; GO:0016021; C:Integral to membrane; ISS.  
DR GO; GO:0004190; F:Aspartic-type endopeptidase activity; ISS.  
DR GO; GO:0050435; P:beta-amyloid metabolism; ISS.  
DR GO; GO:0006509; P:membrane protein ectodomain proteolysis; ISS.  
DR InterPro: IPR001461; Peptidase A1.  
DR InterPro: IPR009119; Pept\_A1\_BACE.  
DR InterPro: IPR009120; Pept\_A1\_BACE1.  
DR InterPro: IPR009007; Pept\_Aspartic.  
DR InterPro: IPR001969; Pept\_Asp\_AS.  
DR PRINTS; PRO1816; BACE1.  
DR PRINTS; PRO1815; BACEFAMILY.  
DR PRINTS; PRO0792; PEPsin.  
DR PROSITE; PS00141; ASP PROTEASE; 1.  
DR Aspartyl protease; Hydrolase; Protease.  
SQ SEQUENCE 501 AA; 55823 MW; 768595CF5517EB7 CRC64;

Query Match 100.0%; Score 16; DB 2; Length 501;  
Best Local Similarity 100.0%; Pred. No. 4.2e-08;  
Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 TOHGIRLPLRSGIGGA 16  
DB 22 TOHGIRLPLRSGIGGA 37

RESULT 4  
Q8CAF4 PRELIMINARY; PRT; 467 AA.  
ID Q8CAF4  
AC Q8CAF4  
DT 01-MAR-2003 (TrEMBLrel. 23, Created)  
DT 01-MAR-2003 (TrEMBLrel. 23, Last sequence update)  
DT 01-MAR-2004 (TrEMBLrel. 26, Last annotation update)  
DE Mus musculus 0 day neonate cerebellum cDNA, RIKEN full-length enriched  
DE library, clone: C230037E16 product:beta-site APP cleaving enzyme, full  
DE insert sequence.  
GN Name=BACE1; Synonym=Bace;  
OS Mus musculus (Mouse).  
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
OC NCB1\_TaxID=10090;  
OX NCB1  
RN [1]  
RP SEQUENCE FROM N.A.  
RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
RX MEDLINE=99279253; PubMed=10349636; DOI=10.1016/S0076-6879(99)03004-9;

RA Carninci P., Hayashizaki Y.;  
 RT "High-efficiency full-length cDNA cloning";  
 RL Meth. Enzymol. 303:19-44(1999).  
 RN (2)  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RX MEDLINE=21085660; PubMed=11217851; DOI=10.1038/35055500;  
 RA RIKEN PANTOM Consortium;  
 RT "Functional annotation of a full-length mouse cDNA collection.";  
 RL Nature 409:685-690(2001).  
 RN [3]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RA The FANTOM Consortium;  
 RT "Analysis of the mouse transcriptome based on functional annotation of  
 RL Nature 420:563-573(2002).  
 RN [4]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RX MEDLINE=20499374; PubMed=11042159; DOI=10.1101/gr.145100;  
 RA Carninci P., Shibata Y., Hayatsu N., Sugahara Y., Shibata K., Itoh M.,  
 RT Kono H., Okazaki Y., Muramatsu M., Hayashizaki Y.;  
 RL "Normalization and subcloning of cap-trapper-selected cDNAs to  
 prepare full-length cDNA libraries for rapid discovery of new genes.";  
 RN Genom. Res. 10:1617-1630(2000).  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RX MEDLINE=20530913; PubMed=11076861; DOI=10.1101/gr.152600;  
 RA Shibata K., Itoh M., Aizawa K., Nagaoka S., Sasaki N., Carninci P.,  
 RT Kono H., Akiyama Y., Nishi K., Kikunishi T., Tashiro H., Itoh M.,  
 RA Sumi N., Ishii Y., Nakamura S., Hazama M., Nishino T., Harada A.,  
 RA Yamamoto R., Matsumoto H., Sakauchi S., Ikegami T., Kashiwagi K.,  
 RA Fujiwaka S., Inoue K., Togawa K., Izawa M., Ohara E., Watanabe M.,  
 RA Yoneda Y., Ishikawa T., Ozawa K., Tanaka T., Matsura S., Kawai J.,  
 RA Okazaki Y., Muramatsu M., Inoue Y., Kira A., Hayashizaki Y.;  
 RT "RIKEN integrated sequence analysis (RISA) system-384-format  
 RL sequencing pipeline with 384 multicapillary sequencer.";  
 RN Genom. Res. 10:1157-1771(2000).  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RA Adachi J., Aizawa K., Akimura T., Arakawa T., Bono H., Carninci P.,  
 RA Fukuda S., Furuno M., Hanagaki T., Hara A., Hashizume W.,  
 RA Hayashida K., Hayatsu N., Hiramoto K., Hiraoka T., Hirozane T.,  
 RA Hori F., Imotani K., Ishii Y., Itoh M., Kagawa I., Kasukawa T.,  
 RA Katoh H., Kawai J., Kojima Y., Kondo S., Kono H., Kouda M., Koya S.,  
 RA Kurihara C., Matsuyama T., Miyazaki R., Murata M., Nakamura M.,  
 RA Niishi K., Nomura K., Numazaki R., Ohno M., Ohsato N., Okazaki Y.,  
 RA Saito R., Satoh H., Sakai C., Sakai K., Sakazume N., Sano H.,  
 RA Sasaki D., Shibata K., Shingawa A., Shiraki T., Sogabe Y., Tagami M.,  
 RA Tagawa A., Takahashi F., Takaku-Akahira S., Takeda Y., Tanaka T.,  
 RA Tomaru A., Toya T., Yasunishi A., Muramatsu M., Hayashizaki Y.;  
 RL Submitted (Apr-2002) to the EMBL/GenBank/DBJ databases.  
 CC -1-SIMILARITY: Belongs to peptidase family A1.  
 DR EMBL; AK082317; BAC38462.1; -  
 DR HSSP; P56817; IFKN.  
 DR MGD; MGI:1346542; Bace1.  
 DR GO; GO:0005768; C:cytosol; ISS.  
 DR GO; GO:0005794; C:extracellular space; TAS.  
 DR GO; GO:0016021; C:integral to membrane; ISS.  
 DR GO; GO:0004190; F:aspartic-type endopeptidase activity; ISS.  
 DR GO; GO:0050435; F:beta-amyloid metabolism; ISS.  
 DR GO; GO:0006509; P:membrane protein ectodomain proteolysis; ISS.  
 DR InterPro; IPR001461; Peptidase A1.  
 DR InterPro; IPR009119; Pept\_A1\_BACE.  
 DR InterPro; IPR009120; Pept\_A1\_BACE1.  
 DR InterPro; IPR009007; Pept\_Aspartic.  
 DR InterPro; IPR001969; Pept\_Asp\_AS.  
 DR PRINTS; PRO1816; BACE1.

DR PRINTS; PRO1815; BACEFAMILY.  
 DR PRINTS; PRO0792; PEPIN.  
 DR PROSITE; PS00141; ASP\_PROTEASE; 1.  
 KW Aspartyl protease; Hydrolase; Protease.  
 SQ SEQUENCE 467 AA; 52063 MW; 31AB674FF1843652 CRC64;  
 Query Match 62.5%; Score 10; DB 2; Length 467;  
 Best local similarity 100.0%; Pred. No. 0.05;  
 Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 QY 4 GIRLPRLSGL 13  
 DB 25 GIRLPRLSGL 34  
 RESULT 5  
 BAE1\_MOUSE  
 ID BAE1\_MOUSE STANDARD; PRT; 501 AA.  
 AC P56818;  
 DT 30-MAY-2000 (Rel. 39, Created)  
 DT 16-OCT-2001 (Rel. 40, Last sequence update)  
 DT 25-OCT-2004 (Rel. 45, Last annotation update)  
 DE Beta-secretase 1 precursor (BC 3.4.23.46) (Beta-site APP cleaving  
 DE enzyme 1) (Beta-site amyloid precursor protein cleaving enzyme 1)  
 DE (Aspartyl protease 2) (Aap 2) (ASP2) (Membrane-associated aspartic  
 DE protease 2) (Memapsin-2).  
 GN Name=BACE1; Synonyms=Bace;  
 OS Mus musculus (Mouse).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 CC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
 OX NCBI\_TaxID=10090;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC MEDLINE=20022972; PubMed=10531052; DOI=10.1126/science.286.5440.735;  
 RA Vassar R., Bennett B.D., Babu-Khan S., Kahn S., Mendiaz E.A.,  
 RA Denis P., Teplow D.B., Ross S., Amarante P., Loebler R., Luo Y.,  
 RA Fisher S., Fuller J., Edwards S., Lille J., Jarosinski M.A.,  
 RA Biere A.L., Curran E., Burgess T., Louis J.-C., Collins F.,  
 RA Treanor D., Rogers G., Citron M.;  
 RT "Beta-secretase cleavage of Alzheimer's amyloid precursor protein by  
 RL the transmembrane aspartic protease BACE.";  
 RN Science 286:735-741(1999).  
 RP REVISIONS TO 6 AND 81-87.  
 RP Bennett B.D., Vassar R., Citron M.;  
 RL Submitted (Jan-2000) to the EMBL/GenBank/DBJ databases.  
 RN [3]  
 RP SEQUENCE FROM N.A.  
 RC MEDLINE=20057170; PubMed=10591213; DOI=10.1038/990107;  
 RA Yan R., Bienkowski M.J., Shuck M.E., Miao H., Toriy M.C., Paulay A.M.,  
 RA Braehler J.R., Strattan N.C., Mathews W.R., Buhl A.E., Carter D.B.,  
 RA Tomasselli A.G., Parodi L.A., Heinrichson R.L., Gurney M.E.;  
 RT "Membrane-anchored aspartyl protease with Alzheimer's disease beta-  
 RT secretase activity.";  
 RL Nature 402:533-537(1999).  
 RN [4]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Head;  
 RX MEDLINE=22354683; PubMed=12466851; DOI=10.1038/nature01266;  
 RA Okazaki Y., Furuno M., Kasukawa T., Adachi J., Bono H., Kondo S.,  
 RA Nishida I., Osato N., Saito R., Suzuki H., Yamana H., Kiyosawa H.,  
 RA Yagi K., Tomaru Y., Hasegawa Y., Nogami A., Schonbach C., Gotohori T.,  
 RA Baldarelli R., Hill D.P., Bult C., Hume D.A., Quackenbush J.,  
 RA Schmitt L.M., Kanapin A., Matsuda H., Batilov S., Beisel K.W.,  
 RA Blake J.A., Bradt D., Brusic V., Chochina C., Corbani L.E., Cousins S.,  
 RA Dalla B., Dragani T.A., Fletcher C.F., Forrest A., Fraser K.S.,  
 RA Grimmerland T., Gariboldi M., Gissi C., Godzik A., Jarvis E.D.,  
 RA Grimmond S., Gustincich S., Hirokawa N., Jackson I.J., Jarvis E.D.,  
 RA Kanai A., Kawai H., Kawasawa Y., Kedzierski R.M., King B.L.,  
 RA Kanagaya A., Kuwajochin I.V., Lee Y., Lenhard B., Lyons P.A.,  
 RA Magloict D.R., Maltais L., Marchionni L., McKenzie L., Mikki H.,  
 RA Nagashima T., Numata K., Okido T., Pavan W.J., Petrea G., Pesole G.,  
 RA Petrovsky N., Pillai R., Pontius J.U., Qi D., Ramachandran S.,

RA Ravasi T., Reed J.C., Reid D.J., Reid J., Ring B.Z., Ringwald M.,  
RA Sandelin A., Schneider C., Sempke C.A., Setou R.D., Shimada K.,  
RA Sultana R., Takenaka Y., Taylor M.S., Teasdale R., Tomita M.,  
RA Verardo R., Wagner L., Wahlestedt C., Wang Y., Watanabe Y., Wells C.,  
RA Wu L.G., Wynshaw-Boris A., Yanagisawa M., Yang L., Yang L.,  
RA Yuan Z., Zeviani M., Zhu Y., Zimmer A., Carninci P., Hayatsu N.,  
RA Hirozane-Kishikawa T., Kono H., Nakamura M., Sakazume N., Sato K.,  
RA Shitaki T., Waki K., Kawai J., Aizawa K., Arakawa T., Fukuda S.,  
RA Hara A., Hashizume W., Imotani K., Ishii Y., Itoh M., Kagawa I.,  
RA Miyazaki A., Sakai K., Sasaki D., Shibata K., Shingawa A.,  
RA Yasunishi A., Yoshino M., Waterston R., Lander E.S., Rogers J.,  
RA Birney E., Hayashizaki Y.,  
RT "Analysis of the mouse transcriptome based on functional annotation of  
RT 60,770 full-length cDNAs";  
RT Nature 420:563-573(2002).  
RN (5)  
RP SEQUENCE FROM N.A.  
RC STRAIN=C57BL/6; TISSUE=Brain;  
RX MEDLINE=22386257; PubMed=12477932; DOI=10.1073/pnas.242603899;  
RA Strausberg R.L., Feingold E.A., Grouse L.H., Derge J.G.,  
RA Klausner R.D., Collins F.S., Wagner L., Shenmen C.M., Schuler G.D.,  
RA Altschul S.F., Zeeberg B., Buetow K.H., Scheffer C.F., Bhat N.K.,  
RA Hopkins R.F., Jordan H., Moore T., Max S.I., Wang J., Hsieh F.,  
RA Diatchenko L., Marusina K., Farmer A.A., Rubin G.M., Hong L.,  
RA Stopleton M., Soares M.B., Bonaldo M.F., Casavant T.L., Scheetz T.E.,  
RA Brownstein M.J., Ueda T.B., Toshiyuki S., Carninci P., Prange C.,  
RA Raza S.S., Loquellano N.A., Peters G.J., Abramson R.D., Mallary S.J.,  
RA Bosak S.A., McEwan P.J., McKernan K.J., Malek J.A., Gunaratne P.H.,  
RA Richards S., Worley K.C., Hale S., Garcia A.M., Gay L.J., Hulik S.W.,  
RA Villalón D.K., Muzny D.M., Sodergren E.J., Lu X., Gibbs R.A.,  
RA Fahy J., Helton E., Ketterman M., Madan A.C., Rodriques S., Sanchez A.,  
RA Whiting M., Madan A., Young A.C., Shevchenko Y., Bouffard G.G.,  
RA Blakesley R.W., Touchman J.W., Green E.D., Dickson M.C.,  
RA Rodriguez A.C., Grimwood J., Schmutz J., Myers R.M.,  
RA Butterfield Y.S.N., Krzywinski M.I., Skalska U., Smalins D.E.,  
RA Scherch A., Schein J.E., Jones S.J.M., Marra M.A.,  
RT "Generation and initial analysis of more than 15,000 full-length human  
RT and mouse cDNA sequences";  
RT Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903(2002).  
CC -1- FUNCTION: Responsible for the proteolytic processing of the  
CC amyloid precursor protein (APP). Cleaves at the amino terminus of  
CC the A-beta peptide sequence, between residues 671 and 672 of APP,  
CC leads to the generation and extracellular release of beta-cleaved  
CC soluble APP, and a corresponding cell-associated carboxy-terminal  
CC fragment which is later released by gamma-secretase (By  
CC similarity).  
CC -1- CATALYTIC ACTIVITY: Broad endopeptidase specificity. Cleaves Glu-  
CC Val-Asn-Ileu-|-Asp-Ala-Glu-Phe in the Swedish variant of  
CC Alzheimer's amyloid precursor protein.  
CC -1- SUBCELLULAR LOCATION: Type I membrane protein.  
CC -1- TISSUE SPECIFICITY: Brain.  
CC -1- SIMILARITY: Belongs to the peptidase A1 family.  
CC -----  
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CC or send an email to [license@isb-sib.ch](mailto:license@isb-sib.ch)).  
CC -----  
CC EMBL: AF190726; AAF04143.2; -;  
CC EMBL: AF200346; AAF17082.1; -;  
CC EMBL: AK014464; BAB29370.1; -;  
CC EMBL: BC048189; AAH48189.1; -;  
CC HSSP: P56817; 1M4H.  
CC MEROPS: A01.004; -;  
CC MGD: MGI:1346542; Bace1.  
CC InterPro: IPR009119; Pept\_A1\_BACE.  
CC InterPro: IPR009120; Pept\_A1\_BACE1.  
CC InterPro: IPR001969; Pept\_A1\_AS.  
CC InterPro: IPR009007; Pept\_Aspartic.  
CC InterPro: IPR001461; Peptidase\_A1.

DR Pfam: PF00026; Asp; 1.  
DR PRINTS: PR01816; BACE1.  
DR PRINTS: PR01815; BACEFAMILY.  
DR PRINTS: PR00792; PEPsin.  
DR PROSITE: PS00141; ASP-PROTEASE, 1.  
KW Aspartyl protease; Glycoprotein; Hydrolase; Signal; Transmembrane;  
KW Zymogen.  
FT SIGNAL 1 21 Potential.  
FT PROPEP 22 45 Potential.  
FT CHAIN 46 501 Beta-secretase 1.  
FT DOMAIN 22 457 Extracellular (Potential).  
FT TRANSMEM 458 478 Potential.  
FT DOMAIN 479 501 Cytoplasmic (Potential).  
FT ACT\_SITE 93 93 By similarity.  
FT ACT\_SITE 289 289 By similarity.  
FT DISULFID 216 420 By similarity.  
FT DISULFID 278 443 By similarity.  
FT DISULFID 330 380 By similarity.  
FT CARBOHYD 153 153 N-linked (GlcNAc...) (Potential).  
FT CARBOHYD 172 172 N-linked (GlcNAc...) (Potential).  
FT CARBOHYD 223 223 N-linked (GlcNAc...) (Potential).  
FT CARBOHYD 354 354 N-linked (GlcNAc...) (Potential).  
SQ SEQUENCE 501 AA; 55747 MW; C085A01315E474E CRC64;  
Query Match 62.5%; Score 10; DB 1; Length 501;  
Best Local Similarity 100.0%; Pred. No. 0.053;  
Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
QY 4 GIRLPRLSGL 13  
DB 25 GIRLPRLSGL 34  
RESULT 6  
BAEL RAT STANDARD; PRT; 501 AA.  
AC P56819;  
DT 30-MAY-2000 (Rel. 39, Last sequence update)  
DT 30-MAY-2000 (Rel. 39, Last annotation update)  
DT 25-OCT-2004 (Rel. 45, Last annotation update)  
DE beta-secretase 1 precursor (EC 3.4.23.46) (Beta-site APP cleaving  
DE enzyme 1) (Beta-site amyloid precursor protein cleaving enzyme 1)  
DE (Aspartyl) protease 2) (Asp 2) (ASP2) (Membrane-associated aspartic  
DE protease 2) (Memapsin-2).  
GN Name=Bace1; Synonyms=Bace;  
OS Rattus norvegicus (Rat).  
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Rattus.  
OX NCBI\_TaxID=10116;  
RN (1)  
RP SEQUENCE FROM N.A.  
RX MEDLINE=20002972; PubMed=10531052; DOI=10.1126/science.286.5440.735;  
RA Vassar R., Bennett B.D., Babu-Khan S., Kahn S., Mendiaz E.A.,  
RA Denis P., Teplow D.B., Ross S., Amarante P., Loefler R., Luo Y.,  
RA Fisher S., Fuller J., Edenson S., Lile J., Jarosinski M.A.,  
RA Biere A.L., Curran E., Burgess T., Louis J.-C., Collins F.,  
RA Treanor J., Rogers G., Citron M.,  
RT "Beta-secretase cleavage of Alzheimer's amyloid precursor protein by  
RT the transmembrane aspartic protease BACE";  
RT Science 286:735-741(1999).  
RL -1- FUNCTION: Responsible for the proteolytic processing of the  
CC amyloid precursor protein (APP). Cleaves at the amino terminus of  
CC the A-beta peptide sequence, between residues 671 and 672 of APP,  
CC leads to the generation and extracellular release of beta-cleaved  
CC soluble APP, and a corresponding cell-associated carboxy-terminal  
CC fragment which is later released by gamma-secretase (By  
CC similarity).  
CC -1- CATALYTIC ACTIVITY: Broad endopeptidase specificity. Cleaves Glu-  
CC Val-Asn-Ileu-|-Asp-Ala-Glu-Phe in the Swedish variant of  
CC Alzheimer's amyloid precursor protein.  
CC -1- SUBCELLULAR LOCATION: Type I membrane protein.  
CC -1- SIMILARITY: Belongs to the peptidase A1 family.  
CC -----



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DR EMBL; AF190727; AF04144.1; -  
 DR HSSP; P56817; IM4H.  
 DR MEROPS; A01.004; -  
 DR RGD; 2191; Bace.  
 DR InterPro; IPR009119; Pept\_A1\_BACE.  
 DR InterPro; IPR009120; Pept\_A1\_BACE1.  
 DR InterPro; IPR001969; Pept\_Asp\_AS.  
 DR InterPro; IPR009007; Pept\_Aspartic.  
 DR InterPro; IPR001461; Peptidase\_A1.  
 DR Pfam; PF00026; Asp\_1.  
 DR PRINTS; PRO1816; BACE1.  
 DR PRINTS; PRO1815; BACEFAMILY.  
 DR PROSITE; PRO0792; PEPSIN.  
 DR PROSITE; PS00141; ASP\_PROTEASE; 1.  
 KW Aspartyl protease; Glycoprotein; Hydrolase; Signal; Transmembrane;  
 FT SIGINT; 1 21  
 FT PROPEP 22 45  
 FT CHAIN 46 501  
 FT DOMAIN 22 457  
 FT TRANSMEM 458 478  
 FT DOMAIN 479 501  
 FT ACT SITE 93 93  
 FT ACT SITE 289 289  
 FT DISULFID 216 420  
 FT DISULFID 278 443  
 FT DISULFID 310 380  
 FT CARBOHYD 153 153  
 FT CARBOHYD 172 172  
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Query Match  
 Best Local Similarity 100.0%; Pred. No. 0.053; Length 501;  
 Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Oy 4 GIRLPLRSGL 13  
 Db 25 GIRLPLRSGL 34

RESULT 7  
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 AC O8BOY4  
 DT 01-MAR-2003 (TREMBLrel. 23. Last sequence update)  
 DT 01-MAR-2003 (TREMBLrel. 23. Last sequence update)  
 DE Mus musculus adult male corpora quadrigemina CDNA, RIKEN full-length  
 DE enriched library, clone:BS30346M13 product:beta-site APP cleaving  
 GN Name=Bace1; Synonyms=Bace;  
 OS Mus musculus (Mouse).  
 OC Eukaryota; Metazoa; Chordata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
 OX NCBI\_TaxId=10090;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RX MEDLINE=99279253; PubMed=10349636; DOI=10.1016/S0076-6879(99)03004-9;  
 RA Carninci P., Hayashizaki Y.;  
 RT "High-efficiency full-length cDNA cloning.";  
 RN Meth. Enzymol. 303:19-44(1999). [2]

RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RX MEDLINE=21085660; PubMed=11217851; DOI=10.1038/35055500;  
 RA RIKEN FANTOM Consortium;  
 RT "Functional annotation of a full-length mouse cDNA collection.";  
 RL Nature 409:685-690(2001).  
 RN [3]

RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RX MEDLINE=20499374; PubMed=11042159; DOI=10.1101/gr.145100;  
 RA Carninci P., Shibata Y., Hayatsu N., Sugahara Y., Shibata K., Itoh M.,  
 RT "Normalization and subcloning of cap-trapper-selected cDNAs to  
 RT prepare full-length cDNA libraries for rapid discovery of new genes.";  
 RL Genome Res. 10:1617-1630(2000).  
 RN [5]

RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RX MEDLINE=20530913; PubMed=11076861; DOI=10.1101/gr.152600;  
 RA Shibata K., Itoh M., Aizawa K., Nagaoka S., Sasaki N., Carninci P.,  
 RA Kono H., Akiyama U., Nishi K., Kitsuana T., Tashiro H., Itoh M.,  
 RA Sumi N., Ishii Y., Nakamura S., Hazama M., Nishine T., Harada A.,  
 RA Yamamoto R., Matsumoto H., Sakaguchi S., Ikegami T., Kashiwagi K.,  
 RA Fujiwaki S., Inoue K., Ozawa Y., Izawa M., Ohara E., Wataniki M.,  
 RA Yoneda Y., Ishikawa T., Ozawa K., Tanaka T., Matsumura S., Kawai J.,  
 RA Okazaki Y., Muramatsu M., Inoue Y., Kira A., Hayashizaki Y.;  
 RT RIKEN integrated sequence analysis (RISA) system-384-format  
 RT sequencing pipeline with 384 multicapillary sequencer.";  
 RL Genome Res. 10:1757-1771(2000).  
 RN [6]

RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RA Adachi U., Aizawa K., Akimura T., Arakawa T., Bono H., Carninci P.,  
 RA Fukuda S., Furuno M., Hanagaki T., Hara A., Hashizume W.,  
 RA Hayashida K., Hayatsu N., Hiramoto K., Hirooka T., Hirozane T.,  
 RA Hori F., Imocani K., Ishii Y., Itoh M., Kagawa I., Kasukawa T.,  
 RA Katoh H., Kawai J., Kojima Y., Kondo S., Kono H., Kouda M., Koya S.,  
 RA Kurihara C., Matsuyama T., Miyazaki A., Murata M., Nakamura M.,  
 RA Nishi K., Nomura K., Numazaki R., Ohno M., Ohsato N., Okazaki Y.,  
 RA Saito K., Saiton H., Sakai C., Sakai K., Sakazume N., Sano H.,  
 RA Tagawa A., Takahashi P., Shingawa A., Shiraki T., Sogabe Y., Tagami M.,  
 RA Tomaru A., Toya T., Yasunishi A., Muramatsu M., Hayashizaki Y.;  
 RL Submitted (JUL-2001) to the EMBL/Genbank/DBJ databases.  
 CC -1- SIMILARITY: Belongs to the EMBL/Genbank/DBJ databases.  
 DR EMBL; AK046175; BAC32620.1; -  
 DR HSSP; P56817; 1FKN

DR MGD; MGI:1346542; Bace1.  
 DR GO; GO:0005768; C:cytosome; ISS.  
 DR GO; GO:0005615; C:extracellular space; TAS.  
 DR GO; GO:0005794; C:Golgi apparatus; ISS.  
 DR GO; GO:0016021; C:integral to membrane; ISS.  
 DR GO; GO:0004190; F:aspartic-type endopeptidase activity; ISS.  
 DR GO; GO:0050435; F:beta-amyloid metabolism; ISS.  
 DR GO; GO:0006509; P:membrane protein ectodomain proteolysis; ISS.  
 DR InterPro; IPR001461; Peptidase\_A1.  
 DR InterPro; IPR009119; Pept\_A1\_BACE.  
 DR InterPro; IPR009120; Pept\_A1\_BACE1.  
 DR InterPro; IPR009007; Pept\_Aspartic.  
 DR InterPro; IPR001969; Pept\_Asp\_AS.  
 DR PRINTS; PRO1816; BACE1.  
 DR PRINTS; PRO1815; BACEFAMILY.  
 DR PROSITE; PRO0792; PEPSIN  
 DR PROSITE; PS00141; ASP\_PROTEASE; 1.  
 KW Aspartyl protease; Hydrolase; Protease.



RC STRANL-C57BL/6J; TISSUE=Spinal cord;  
RA Adachi J., Aizawa K., Akimura T., Arakawa T., Bono H., Carninci P.,  
RA Fukuda S., Furuno M., Hanagaki T., Hara A., Hashizume W.,  
RA Hayashida K., Hayatsu N., Hiramoto K., Hirooka T., Hirozane T.,  
RA Horii F., Imclani K., Ishii Y., Itoh M., Kagawa I., Kasukawa T.,  
RA Katoh H., Kawai J., Kojima Y., Kondo S., Konno H., Kouda M., Koya S.,  
RA Kurihara C., Matsuyama T., Miyazaki A., Murata M., Nakamura M.,  
RA Nishi K., Nomura K., Numasaki R., Ohno M., Ohsato N., Okazaki Y.,  
RA Saeto R., Satoh H., Sakai C., Sakai K., Sakazume N., Sano H.,  
RA Sawaki D., Shibata K., Shinigawa A., Shiraki T., Sogabe Y., Tagami M.,  
RA Tagawa A., Takahashi F., Takeku-Akaiwa S., Takeeda Y., Tanaka T.,  
RA Tomaru A., Toyota T., Yasunishi A., Yamamatsu M., Hayashiaki Y.;  
CC Submitted (JUL-2001) to the EMBL/GenBank/DBJ databases.  
RL -1- SIMILARITY: Belongs to peptidase family A1.  
DR EMBL; AK049626; BAC33844.1; -.  
DR HSSP; P56817; 1FKN.  
DR MGD; MG1:1346542; Bacel.  
DR GO; GO:0005768; C:endosome; ISS.  
DR GO; GO:0005615; C:extracellular space; TAS.  
DR GO; GO:0005794; C:Golgi apparatus; ISS.  
DR GO; GO:0016021; C:integral to membrane; ISS.  
DR GO; GO:001490; F:aspartic-type endopeptidase activity; ISS.  
DR GO; GO:0005035; P:beta-amylid metabolism; ISS.  
DR GO; GO:0006049; P:membrane protein ectodomain proteolysis; ISS.  
DR InterPro: IPRO01461; Peptidase A1.  
DR InterPro: IPRO09119; Pept\_A1\_BACE.  
DR InterPro: IPRO09120; Pept\_A1\_BACE.  
DR InterPro: IPRO09007; Pept\_AspArtic.  
DR InterPro: IPRO01969; Pept\_Asp\_AS.  
DR PRINTS; PRO1816; BACEFAMILY.  
DR PRINTS; PRO1815; BACEFAMILY.  
DR PRINTS; PRO0792; PEPSIN.  
DR PROSITE; PS00141; ASP\_PROTEASE; 1.  
KW Aspartyl protease; Hydrolase; Protease.  
SQ SEQUENCE 501 AA; 55761 MW; B41DDA8B64647663 CRC64;

QY 4 GIRLPURSGL 13  
|||||  
Db 25 GIRLPURSGL 34

Query March 62.5%; Score 10; DB 2; Length 501;  
Best Local Similarity 100.0%; Pred. No. 0.053;  
Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0.

RESULT 9  
Q7SZ18 PRELIMINARY; PRT; 814 AA.  
ID Q7SZ18  
AC Q7SZ18;  
DT 01-OCT-2003 (TrEMBLrel. 25, Created)  
DT 01-OCT-2003 (TrEMBLrel. 25, Last sequence update)  
DE 01-MAR-2004 (TrEMBLrel. 26, Last annotation update)  
DE SWI1 alpha (Fragment).  
OS Oryzias latipes (Medaka fish) (Japanese ricefish).  
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
OC Actinopterygii; Neopterygii; Teleostei; Euteleostei; Neoteleostei;  
OC Acantbomorphes; Acanthopterygii; Percomorpha; Atherinomorpha;  
OC Belontiiformes; Atherinchthyidae; Oryziinae; Oryzias.  
OX NCBI\_Taxid=8090;  
RN [1]  
RS SEQUENCE FROM N.A.  
RC TISSUE=Tectis;  
RX MEDLINE=22660315; PubMed=12759374;  
RA Lee J., Iwai T., Yokota T., Yamashita M.;  
RT "Temporally and spatially selective loss of Rec8 protein from meiotic chromosomes during mammalian meiosis";  
RL J. Cell Sci. 116:2781-2790(2003).  
CC -1- SIMILARITY: Belongs to the ABC transporter family.  
DR EMBL; AB097255; BAC76893.1; -.  
DR HSSP; Q9XOR4; 1B69.  
DR GO; GO:0016020; C:membrane; IEA.  
DR GO; GO:0005524; F:ATP binding; IEA.  
DR GO; GO:0042626; F:ATPase activity, coupled to transmembrane m. . ; IEA.

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DR GO: GO:0007059; P:chromosome segregation; IEA.
DR GO: GO:0006810; P:transport; IEA.
DR InterPro: IPR003439; ABC transporter.
DR InterPro: IPR003405; SMC_C.
DR InterPro: IPR010935; SMC_hinge.
DR Pfam: PF02483; SMC_C; 1.
DR Pfam: PF06470; SMC_hinge; 1.
DR Prodom: PD000006; ABC transporter; 1.
DR Prosite: PS00211; ABC_TRANSPORTER_1; UNKNOWN_1.
KM ATP-binding.
FT NON_TER
SQ SEQUENCE 814 AA; 94132 MW; 8653EC762EC6A5A CRC64;

Query Match
Best Local Similarity 50.0%; Score 8; DB 2; Length 814;
Matches 8; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 5 IRLPLRSG 12
Db 521 IRLPLRSG 528

RESULT 10
Q9JPF6 PRELIMINARY; PRT; 97 AA.
ID Q9JPF6
AC Q9JPF6
DT 01-OCT-2000 (TRENBLrel. 15, Created)
DT 01-OCT-2000 (TRENBLrel. 15, Last sequence update)
DT 01-OCT-2003 (TRENBLrel. 25, Last annotation update)
DE Hypothetical protein rch9.
GN Name=rch9;
OS Neisseria meningitidis.
OC Bacteria; Proteobacteria; Betaproteobacteria; Neisseriales;
OC Neisseriaceae; Neisseria.
OX NCBI_TaxID=487;
RN [1]
RP SEQUENCE FROM N.A.
RC STRAIN=Z2491;
RX DOI=10.1128/JAI.68.4.2082-2095.2000;
RA Klee S.R., Nassif X., Kusecek B., Werker P., Beretti J.L., Achtman M.,
RA Tinsley C.R.;
RT "Molecular and biological analysis of eight genetic islands that
RT distinguish Neisseria meningitidis from the closely related pathogen
RT Neisseria gonorrhoeae".
RL Infect. Immun. 68:2082-2095(2000).
DR EMBL; AJ391256; CAB71967.1; -.
KM Hypothetical protein.
SQ SEQUENCE 97 AA; 10716 MW; 7EDF863F7B6531F9 CRC64;

Query Match
Best Local Similarity 43.8%; Score 7; DB 2; Length 97;
Matches 7; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 9 LRSLGIG 15
Db 81 LRSLGIG 87

RESULT 11
Q9JTC5 PRELIMINARY; PRT; 125 AA.
ID Q9JTC5
AC Q9JTC5
DT 01-OCT-2000 (TRENBLrel. 15, Created)
DT 01-OCT-2000 (TRENBLrel. 15, Last sequence update)
DT 01-JUN-2003 (TRENBLrel. 24, Last annotation update)
DE Hypothetical protein NMA1874.
GN OrderedlocusNames=NMA1874;
OS Neisseria meningitidis (serogroup A).
OC Bacteria; Proteobacteria; Betaproteobacteria; Neisseriales;
OC Neisseriaceae; Neisseria.
OX NCBI_TaxID=65699;
RN [1]
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RP SEQUENCE FROM N.A.
RC STRAIN=Z2491 / Serogroup A / Serotype 4A;
RX MEDLINE=20222556; PubMed=10761919; DOI=10.1038/35006655;
RA Parthill J., Achtman M., James K.D., Bentley S.D., Churcher C.M.,
RA Klee S.R., Morrelli G., Baaham D., Brown D., Chillingworth T.,
RA Davies R.M., Davis P., Devlin K., Fellwell T., Hamlin N., Holroyd S.,
RA Jajels K., Leather S., Mould S., Mungall K.L., Quail M.A.,
RA Rajandream M.A., Rutherford K.M., Simmonds M., Skelton J.,
RA Whitehead S., Spratt B.G., Barrall B.G.;
RT "Complete DNA sequence of a serogroup A strain of Neisseria
RT meningitidis Z2491."
RL Nature 404:502-506(2000).
DR EMBL; AL162757; CAB85097.1; -.
DR PIR; E81814; E81814.
KM Complete proteome; Hypothetical protein.
SQ SEQUENCE 125 AA; 13950 MW; 5B9C7D782E89884D CRC64;

Query Match
Best Local Similarity 43.8%; Score 7; DB 2; Length 125;
Matches 7; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 9 LRSLGIG 15
Db 109 LRSLGIG 115

RESULT 12
Q7NTK4 PRELIMINARY; PRT; 181 AA.
ID Q7NTK4
AC Q7NTK4
DT 01-MAR-2004 (TRENBLrel. 26, Created)
DT 01-MAR-2004 (TRENBLrel. 26, Last sequence update)
DT 01-MAR-2004 (TRENBLrel. 26, Last annotation update)
DE Cytochrome b561
GN Name=cybB; OrderedlocusNames=CV3050;
OS Chromobacterium violaceum.
OC Bacteria; Proteobacteria; Betaproteobacteria; Neisseriales;
OC Neisseriaceae; Chromobacterium.
OX NCBI_TaxID=536;
RN [1]
RP SEQUENCE FROM N.A.
RC STRAIN=ATCC 12472 / DSM 30191;
RX MEDLINE=22882880; PubMed=14500782; DOI=10.1073/pnas.1832124100;
RA Vasconcelos A.T.R., de Almeida D.F., Hungria M., Gularaes C.T.,
RA Antonio R.V., Almeida F.C., de Almeida L.G.P., de Almeida R.,
RA Alves-Gomes J.A., Andrade E.M., Azevedo V., de Araujo M.F.F.,
RA Astolfi-Filho S., Azevedo V., Baptista A.J., Batista L.A.M.,
RA Batista U.S., Belo A., van den Berg C., Bogo M., Bonatto S.,
RA Bordignon J., Brígido M.M., Brito C.A., Brocchi M., Burtly H.A.,
RA Camargo A.A., Cardoso D.D.P., Carneiro N.P., Carreiro D.M.,
RA Carvalho C.M.B., Cascardo J.C.M., Cavada B.S., Chueire L.M.O.,
RA Creczynski-Pasa T.B., Cunha-Junior N.C., Fagundes N., Falcao C.L.,
RA Fancinatti F., Farias I.P., Felipe M.S.S., Ferrari L.P., Ferro J.A.,
RA Ferro M.I.T., Franco G.R., Freitas N.S.A., Furlan L.R.,
RA Gazzinelli R.T., Gomes E.A., Goncalves P.R., Grangelito T.B.,
RA Gracatapaglia D., Girsard E.C., Hanna E.S., Jardim S.N., Laurino J.,
RA Leoi L.C.T., Lima L.F.A., Loureiro M.F., Lyra M.C.C.P.,
RA Madeira H.M.F., Manfro G.P., Maranhao A.O., Martins W.S.,
RA di Mauro S.M.Z., de Medeiros S.R.B., Meisner R.V., Moreira M.A.M.,
RA Nascimento F.F., Nicolas M.F., Oliveira J.G., Oliveira S.C.,
RA Paixao R.F.C., Parente J.A., Pedrosa F.O., Pena S.D.J., Pereira J.O.,
RA Pereira M., Pinto L.S.R.C., Pinto L.S., Porto J.I.R., Potrich D.P.,
RA Ramalho-Neto C.E., Reis A.M.M., Rigo L.U., Rondinelli E.,
RA Santos E.B.P., Santos F.R., Schneider M.P.C., Senanez H.N.,
RA Silva A.M.R., da Silva A.L.C., Silva D.W., Silva R., Simoes I.C.,
RA Simon D., Soares C.M.A., Soares R.B.A., Souza E.M., Souza K.R.L.,
RA Souza R.C., Steffens M.B.R., Steindel M., Teixeira S.R., Urmenyi T.,
RA Vettore A., Wasson R., Zaha A., Simpson A.J.G.;
RT "The complete genome sequence of Chromobacterium violaceum reveals
RT remarkable and exploitable bacterial adaptability".
RL Proc. Natl. Acad. Sci. U.S.A. 100:11660-11665(2003).
DR EMBL; AE016920; AA060719.1; -.
DR InterPro; IPR011577; CybB561_bact.
```

DR Pfam: PF01292; N1\_hydr\_CyTB; 1.  
 KW Complete proteome  
 SQ SEQUENCE 181 AA; 19650 MW; 94FBFB853494F08 CRC64;

Query Match  
 Best Local Similarity 43.8%; Score 7; DB 2; Length 181;  
 Matches 7; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 8 PLRSLG 14  
 DB 42 PLRSLG 48

RESULT 13

Q66F29 PRELIMINARY; PRT; 239 AA.  
 AC Q66F29;  
 DT 25-OCT-2004 (TREMBLrel. 28, Created)  
 DT 25-OCT-2004 (TREMBLrel. 28, Last sequence update)  
 DT 25-OCT-2004 (TREMBLrel. 28, Last annotation update)  
 DE Putative ABC phosphonate transporter, ATP binding protein, also  
 DE putative C-P lyase component.  
 GN Name=phnL; ORFNames=YPT80511;  
 OS Yersinia pseudotuberculosis IP 32953.  
 OC Bacteria; Proteobacteria; Gammaproteobacteria; Enterobacteriales;  
 OC Enterobacteriaceae; Yersinia.  
 OK NCBI\_TaxID=273123;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=IP 32953;  
 RX PubMed=15358958;  
 RA Chain P.S.G., Carmiel E., Larimer F.W., Lamerdin J., Stouland P.O.,  
 RA Regla W.M., Georgescu A.M., Verge L.M., Land M.L., Motin L.V.,  
 RA Brubaker R.R., Fowler J., Hinnelbuch B.J., Marcu M., Medigue C.,  
 RA Simonet M., Chena-Francoise V., Souza B., Dacheux D., Elliott J.M.,  
 RA Derise A., Hauser L.J., Garcia E.;  
 RA "Insights into the genome evolution of Yersinia pestis through whole  
 RA genome comparison with Yersinia pseudotuberculosis";  
 RT Proc. Natl. Acad. Sci. U.S.A. 101:13826-13831(2004).  
 DR EMBL; BX936398; CAH19751.1; -;  
 DR GO; GO:0016829; P:lyase activity; IEA.  
 DR InterPro: IPR003593; AAA ATPase.  
 DR InterPro: IPR003439; ABC\_transporter.  
 DR Pfam; PF00005; ABC\_tran; 1.  
 DR Pfam; PD000006; ABC\_transporter; 1.  
 DR SMART; SM00382; AAA; 1.  
 DR PROSITE; PS00211; ABC\_TRANSPORTER\_1; 1.  
 DR PROSITE; PS00893; ABC\_TRANSPORTER\_2; 1.  
 KW ATP-binding; Lyase.  
 SQ SEQUENCE 239 AA; 26695 MW; 4096220E6760B9FB CRC64;

Query Match  
 Best Local Similarity 43.8%; Score 7; DB 2; Length 239;  
 Matches 7; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 2 QHGIRLP 8  
 DB 20 QHGIRLP 26

RESULT 14

Q82BF4 PRELIMINARY; PRT; 239 AA.  
 AC Q82BF4; Q7CKG6;  
 DT 01-MAR-2002 (TREMBLrel. 20, Created)  
 DT 01-MAR-2002 (TREMBLrel. 20, Last sequence update)  
 DT 25-OCT-2004 (TREMBLrel. 28, Last annotation update)  
 DE Phosphonate ABC transporter.  
 DE Name=phnL; OrderedLocNames=YPO3462, Y0723;  
 GN Yersinia pestis.  
 OS Bacteria; Proteobacteria; Gammaproteobacteria; Enterobacteriales;  
 OC Enterobacteriaceae; Yersinia.

OK NCBI\_TaxID=632;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=CO-92 / Bivovar Orientalis;  
 RX MEDLINE=21470413; PubMed=11586360; DOI=10.1038/35097083;  
 RA Parthill J., Wren B.W., Thomson N.R., Tibball R.W., Holden M.T.G.,  
 RA Prentice M.B., Sebahia M., James K.D., Churcher C.M., Mungall K.L.,  
 RA Baker S., Basham D., Bentley S.D., Brooks K., Cerdano-Tarraga A.-M.,  
 RA Chillingworth T., Cronin A., Davies R.M., Davis P., Dougan G.,  
 RA Fellwell T., Hamlin N., Holroyd S., Jagels K., Karlyshev A.V.,  
 RA Leachner S., Moule S., Oyston P.C.F., Quail M.A., Rutherford K.M.,  
 RA Simmonds M., Skellon J., Stevens K., Whitehead S., Barrett B.G.;  
 RA "Genome sequence of Yersinia pestis, the causative agent of plague";  
 RA Nature 413:523-527(2001).  
 RN [2]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=KIM5 / Bivovar Mediaevalis;  
 RX MEDLINE=22137863; PubMed=12142430;  
 RX DOI=10.1128/JB.184.16.4601-4611.2002;  
 RA Deng W., Burland V., Plunkett G. III, Boutin A., Mayhew G.F., Liss P.,  
 RA Perna N.T., Rose D.J., Mau B., Zhou S., Schwartz D.C.,  
 RA Fetherston J.D., Lindler L.E., Brubaker R.R., Plano G.V.,  
 RA Straley S.C., McDonough K.A., Nilles M.L., Mateon J.S., Blattner F.R.,  
 RA Perry R.D.;  
 RA "Genome sequence of Yersinia pestis KIM";  
 RT J. Bacteriol. 184:4601-4611(2002).  
 CC -1- SIMILARITY: Belongs to the ABC transporter family.  
 DR EMBL; AJ414157; CAC92691.1; -;  
 DR EMBL; AE013674; NAM84311.1; -;  
 DR PIR; AG0420; AG0420.  
 DR GO; GO:0016020; C:membrane; IEA.  
 DR GO; GO:0005524; F:ATP binding; IEA.  
 DR GO; GO:0042625; F:ATPase activity; IEA.  
 DR GO; GO:0000166; F:nucleotide binding; IEA.  
 DR GO; GO:0006810; P:transport; IEA.  
 DR Pfam; PF00005; ABC\_tran; 1.  
 DR Pfam; PD000006; ABC\_transporter; 1.  
 DR SMART; SM00382; AAA; 1.  
 DR PROSITE; PS00211; ABC\_TRANSPORTER\_1; 1.  
 DR PROSITE; PS00893; ABC\_TRANSPORTER\_2; 1.  
 KW ATP-binding; Complete proteome.  
 SQ SEQUENCE 239 AA; 26667 MW; 59AB94CC3760B9FB CRC64;

Query Match  
 Best Local Similarity 43.8%; Score 7; DB 2; Length 239;  
 Matches 7; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 2 QHGIRLP 8  
 DB 20 QHGIRLP 26

RESULT 15

Q9RXU2 PRELIMINARY; PRT; 259 AA.  
 ID Q9RXU2;  
 AC Q9RXU2;  
 DT 01-MAY-2000 (TREMBLrel. 13, Created)  
 DT 01-MAY-2000 (TREMBLrel. 13, Last sequence update)  
 DT 01-JUN-2003 (TREMBLrel. 24, Last annotation update)  
 DE Hypothetical protein DR0214.  
 DE OrderedLocNames=DR0214;  
 GN Deinococcus radiodurans.  
 OS Deinococcus radiodurans.  
 OC Bacteria; Deinococcus-Thermus; Deinococci; Deinococcales;  
 OC Deinococcaceae; Deinococcus.  
 OK NCBI\_TaxID=1299;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=R1 / ATCC 13939 / DSM 20539 / NCIB 9279;  
 RX MEDLINE=20036896; PubMed=10567266; DOI=10.1126/science.286.5444.1571;  
 RA White O., Sisen J.A., Heidelberg J.F., Hickey E.K., Peterson J.D.,  
 RA Dodson R.J., Haft D.H., Gwinn M.L., Nelson W.C., Richardson D.L.,  
 RA Moffat K.S., Qin H., Jiang L., Pamphile W., Crosby M., Shen M.,  
 RA Vamathevan J.J., Lam P., McDonald L.A., Uitterback T.R., Zalewski C.,

RA Makarova K.S., Aravind L., Daly M.J., Minton K.W., Fleischmann R.D.,  
 RA Ketchum K.A., Nelson K.E., Salzberg S.L., Smith H.O., Venter J.C.,  
 RA Fraser C.M.;  
 RT "Genome sequence of the radioresistant bacterium Deinococcus  
 RT radiodurans R1." Science 286:1571-1577(1999).  
 RL EMBL; AE001883; AAF09805.1; -.  
 DR PIR; A75547; A75547.  
 DR TIGR; DR0214; -.  
 KW Complete proteome; Hypothetical protein.  
 SQ SEQUENCE 259 AA; 29103 MM; 49522C0ADD9327CF CRC64;  
  
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 Db 190 HGIRLPL 196

Query Match 43.8%; Score 7; DB 2; Length 259;  
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 Job time : 175 secs

GenCore version 5.1.6  
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OM protein - nucleic search, using frame\_plus\_p2n model

Run on: July 27, 2005, 18:18:13 ; Search time 1937 Seconds  
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400.250 Million cell updates/sec

Title: US-10-726-967A-3

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Searched: 4708233 seqs, 24227607955 residues

Word size: 1

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Minimum DB seq length: 0

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Post-processing: Listing first 45 summaries

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-FGAPEXT=7 -YGAPOP=60 -YGAPEXT=60 -DELDP=6 -DELEXT=7

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7: gb\_ph:\*  
8: gb\_pl:\*  
9: gb\_pr:\*  
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12: gb\_sy:\*  
13: gb\_un:\*  
14: gb\_vl:\*

Pred. No. is the number of results predicted by chance to have a  
score greater than or equal to the score of the result being printed,  
and is derived by analysis of the total score distribution.

#### SUMMARIES

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2	16	100.0	1287	6	AR224122 Sequence
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4	16	100.0	1287	6	AR478808 Sequence

5	16	100.0	1287	6	AR487374	AR487374 Sequence
6	16	100.0	1287	6	AR532014	AR532014 Sequence
7	16	100.0	1287	6	AR540915	AR540915 Sequence
8	16	100.0	1287	6	AR560125	AR560125 Sequence
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10	16	100.0	1287	6	AX573870	AX573870 Sequence
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18	16	100.0	1302	6	AR560106	AR560106 Sequence
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37	16	100.0	1341	6	AR531993	AR531993 Sequence
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#### ALIGNMENTS

RESULT 1  
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LOCUS  
DEFINITION Homo sapiens BACE mRNA for beta-site APP cleaving enzyme isoform  
I-127, complete cds.  
ACCESSION AB089958  
VERSION AB089958.1 GI:34014375  
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SOURCE Homo sapiens (human)  
ORGANISM Homo sapiens  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Primates; Catarrhini; Homiidae; Homo.  
REFERENCE  
AUTHORS Tanahashi, H.  
TITLE A novel alternatively spliced isoform of BACE, I-127 induced by  
cycloheximide treatment  
JOURNAL Unpublished  
AUTHORS 2 (bases 1 to 517)  
TITLE Tanahashi, H.  
DIRECT SUBMISSION  
SUBMITTED (17-AUG-2002) Hiroshi Tanahashi, National Institute of  
Neuroscience, Division of Demyelinating Disease and Aging; 4-1-1  
Ogawabashi, Kodaira, Tokyo 187-8502, Japan  
(E-mail:tanahashicomp.go.jp, Tel:81-042-341-2711(ex.5163),  
Fax:81-042-346-1747)  
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US-10-726-967A-3.01p2n.rge

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/gene34  
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/db\_xref="GI:34014376"  
/translation="MQLPWLMMGAGTGAHGTGHRGLRPLRGLGAPLRLP  
RTRDERGGRGSPFVMDVNLKRSGGGYEWTVGSFPOTNLIVDTSSNFAVG  
AAPPLHRYRQSLSTYRDLRKA"

ORIGIN  
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Score: 1.59e-06  
Percent Similarity: 16.00  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
US-10-726-967A-3 (1-16) x AB089958 (1-517)  
Gaps: 0

ORIGIN  
Alignment Scores:  
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Percent Similarity: 16.00  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
US-10-726-967A-3 (1-16) x AB089958 (1-517)  
Gaps: 0

RESULT 2  
LOCUS AR224122  
DEFINITION Sequence 50 from patent US 6440698. DNA  
ACCESSION AR224122  
VERSION AR224122.1  
KEYWORDS GI:2332782  
SOURCE Unknown.  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1287)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrikson,R.L., Parodi,L.A. and  
TITLE Alzheimer's disease secretase, APP substrates therefor, and uses  
JOURNAL Patent: US 6440698-A 50 27-AUG-2002;  
FEATURES  
source location/Qualifiers  
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/mol\_type="genomic DNA"

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Percent Similarity: 16.00  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
US-10-726-967A-3 (1-16) x AR224122 (1-1287)  
Gaps: 0

ORIGIN  
Alignment Scores:  
Score: 3.27e-06  
Percent Similarity: 16.00  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
US-10-726-967A-3 (1-16) x AR224122 (1-1287)  
Gaps: 0

LOCUS AR269253  
DEFINITION Sequence 50 from patent US 6500667. DNA  
ACCESSION AR269253  
VERSION AR269253.1  
KEYWORDS GI:29700221  
SOURCE Unknown.  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1287)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrikson,R.L., Parodi,L.A. and  
TITLE Aspartyl protease 2 (Asp2) antisense oligonucleotides  
JOURNAL Patent: US 6500667-A 50 31-DEC-2002;  
FEATURES  
source location/Qualifiers  
1..1287  
/mol\_type="genomic DNA"

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Alignment Scores:  
Score: 3.27e-06  
Percent Similarity: 16.00  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
US-10-726-967A-3 (1-16) x AR269253 (1-1287)  
Gaps: 0

RESULT 4  
LOCUS AR478808  
DEFINITION Sequence 50 from patent US 669671. DNA  
ACCESSION AR478808  
VERSION AR478808.1  
KEYWORDS GI:47237528  
SOURCE Unknown.  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1287)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrikson,R.L., Parodi,L.A. and  
TITLE Alzheimer's disease secretase, APP substrates therefor, and uses  
JOURNAL Patent: US 669671-A 50 02-MAR-2004;  
FEATURES  
source location/Qualifiers  
1..1287  
/mol\_type="genomic DNA"

ORIGIN  
Alignment Scores:  
Score: 3.27e-06  
Percent Similarity: 16.00  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
US-10-726-967A-3 (1-16) x AR478808 (1-1287)  
Gaps: 0

ORIGIN  
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Score: 3.27e-06  
Percent Similarity: 16.00  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
US-10-726-967A-3 (1-16) x AR478808 (1-1287)  
Gaps: 0

ACCESSION AR487374  
VERSION AR487374.1 GI:47252472  
KEYWORDS  
SOURCE  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1287)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and  
TITLE Method of identifying agents that inhibit APP processing activity  
JOURNAL Patent: US 6706485-A 50 16-MAR-2004;  
FEATURES  
source  
1..1287  
/organism="unknown"  
/mol\_type="genomic DNA"

ORIGIN

Alignment Scores:  
Pred. No.: 3,27e-06 Length: 1287  
Score: 16.00 Matches: 16  
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Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-3 (1-16) x AR487374 (1-1287)

QY  
1 ThrglnHlgly1leargleuproleuargserglyleuglyg1yala 16  
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Db 64 ACCGACGACGGCATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

RESULT 6  
AR532014  
LOCUS AR532014 1287 bp DNA linear PAT 08-OCT-2004  
DEFINITION Sequence 50 from patent US 6727074.  
ACCESSION AR532014  
VERSION AR532014.1 GI:53920548  
KEYWORDS  
SOURCE Unknown.  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1287)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and  
TITLE Alzheimer's disease secretase, APP substrates therefor, and uses  
therefor  
JOURNAL Patent: US 6727074-A 50 27-APR-2004;  
FEATURES  
source  
1..1287  
/organism="unknown"  
/mol\_type="genomic DNA"

ORIGIN

Alignment Scores:  
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Score: 16.00 Matches: 16  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-3 (1-16) x AR532014 (1-1287)

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1 ThrglnHlgly1leargleuproleuargserglyleuglyg1yala 16  
|||||  
Db 64 ACCGACGACGGCATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

RESULT 7  
AR540915  
LOCUS AR540915 1287 bp DNA linear PAT 08-OCT-2004  
DEFINITION Sequence 50 from patent US 6737510.  
ACCESSION AR540915  
VERSION AR540915.1 GI:53932428

KEYWORDS  
SOURCE Unknown.  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1287)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and  
TITLE Alzheimer's disease secretase, APP substrates therefor, and uses  
thereof  
JOURNAL Patent: US 6737510-A 50 18-MAY-2004;  
FEATURES  
source  
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/organism="unknown"  
/mol\_type="genomic DNA"

ORIGIN

Alignment Scores:  
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DB: 6 Gaps: 0

US-10-726-967A-3 (1-16) x AR540915 (1-1287)

QY  
1 ThrglnHlgly1leargleuproleuargserglyleuglyg1yala 16  
|||||  
Db 64 ACCGACGACGGCATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

RESULT 8  
AR560125  
LOCUS AR560125 1287 bp DNA linear PAT 08-OCT-2004  
DEFINITION Sequence 50 from patent US 6753163.  
ACCESSION AR560125  
VERSION AR560125.1 GI:53970492  
KEYWORDS  
SOURCE Unknown.  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1287)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and  
TITLE Alzheimer's disease secretase, APP substrates therefor, and uses  
therefor  
JOURNAL Patent: US 6753163-A 50 22-JUN-2004;  
FEATURES  
source  
1..1287  
/organism="unknown"  
/mol\_type="genomic DNA"

ORIGIN

Alignment Scores:  
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Score: 16.00 Matches: 16  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-3 (1-16) x AR560125 (1-1287)

QY  
1 ThrglnHlgly1leargleuproleuargserglyleuglyg1yala 16  
|||||  
Db 64 ACCGACGACGGCATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

RESULT 9  
AX105432  
LOCUS AX105432 1287 bp DNA linear PAT 30-APR-2001  
DEFINITION Sequence 50 from Patent WO0123533.  
ACCESSION AX105432  
VERSION AX105432.1 GI:13921541  
KEYWORDS

SOURCE synthetic construct  
ORGANISM other sequences; artificial sequences.  
REFERENCE 1  
AUTHORS Gurney, M. and Bienkowski, M.J.  
TITLE Alzheimer's disease secretase, app substrates therefor, and uses thereof  
JOURNAL Patent: WO 013533-A 50 05-APR-2001;  
Pharmacia & Upjohn Company (US)  
FEATURES  
source 1..1287  
/organism="synthetic construct"  
/mol\_type="unassigned DNA"  
/db\_xref="taxon:32630"  
/note="Hu-Ap2(b) delta TM"

ALIGNMENT Scores:  
Pred. No.: 3.27e-06 Length: 1287  
Score: 16.00 Matches: 16  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-3 (1-16) x AX573870 (1-1287)

QY 1 ThGlnHsglyIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16  
Db 64 ACCCAGCAGCGCATCCGCTGCCCTCGCAGCGCCTCGGCGGCC 111

RESULT 10  
LOCUS AX573870 1287 bp DNA linear PAT 07-JAN-2003  
DEFINITION Sequence 50 from Patent EP1249498.  
ACCESSION AX573870  
VERSION AX573870.1 GI:27551507  
KEYWORDS  
SOURCE synthetic construct  
ORGANISM other sequences; artificial sequences.  
REFERENCE 1  
AUTHORS Gurney, M. and Bienkowski, M.J.  
TITLE Alzheimer's disease secretase, app substrates therefor, and uses thereof  
JOURNAL Patent: EP 1249498-A 50 16-OCT-2002;  
PHARMACIA & UPJOHN COMPANY (US)  
FEATURES  
source 1..1287  
/organism="synthetic construct"  
/mol\_type="unassigned DNA"  
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/note="Hu-Ap2(b) delta TM"

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Pred. No.: 3.27e-06 Length: 1287  
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Query Match: 100.00% Indels: 0  
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US-10-726-967A-3 (1-16) x AX573870 (1-1287)

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Db 64 ACCCAGCAGCGCATCCGCTGCCCTCGCAGCGCCTCGGCGGCC 111

RESULT 11  
LOCUS BD235897 1302 bp DNA linear PAT 17-JUL-2003  
DEFINITION Alzheimer's disease secretase.

ACCESSION BD235897  
VERSION BD235897.1 GI:33045667  
KEYWORDS JP 2002526081-A/13.  
SOURCE Homo sapiens (human)  
ORGANISM Homo sapiens  
REFERENCE 1  
AUTHORS Gurney, M.E., Bienkowski, M.J., Heinrichson, R.L., Parodi, L.A. and Yan, R.  
TITLE Alzheimer's disease secretase  
JOURNAL Patent: JP 2002526081-A 13 20-AUG-2002;  
PHARMACIA AND UPJOHN CO  
COMMENT OS Homo sapiens (human)  
PN JP 2002526081-A/13  
PD 20-AUG-2002  
PF 24-SEP-1999 JP 2000574268  
PR 24-SEP-1998 US 60/101594  
PI MARK E GURNEY, MICHAEL JEROME BIENKOWSKI, ROBERT LEROY PI  
HEINRICHSON, PI  
PI LUIS A. PARODI, RIOIANG YAN  
PC C12N1/09, A61K45/00, A61P25/28, C07K14/47, C07K16/18, C12N1/15, PC  
C12N1/19,  
PC  
C12N1/21, C12N5/10, C12N9/64, C12P21/02, C12P21/08, C12Q1/37, G01N33/15,  
G01N33/50// (C12N1/21, C12R1:19), C12N15/00, C12N5/00 CC  
PC Alzheimer's disease secretase  
FH Key location/Qualifiers  
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/db\_xref="taxon:9606"

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Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-3 (1-16) x BD235897 (1-1302)

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Db 4 ACTCAGCATGGATTCCGCTCCCACTGCGTACCGCGTCTGGGTGCT 51

RESULT 12  
LOCUS AR224103 1302 bp DNA linear PAT 26-SEP-2002  
DEFINITION Sequence 25 from patent US 6440698.  
ACCESSION AR224103  
VERSION AR224103.1 GI:23332763  
KEYWORDS  
SOURCE unknown.  
ORGANISM unknown.  
REFERENCE 1  
AUTHORS Gurney, M.E., Bienkowski, M.J., Heinrichson, R.L., Parodi, L.A. and Yan, R.  
TITLE Alzheimer's disease secretase, APP substrates therefor, and uses thereof  
JOURNAL Patent: US 6440698-A 25 27-AUG-2002;  
FEATURES  
source 1..1302  
/organism="unknown"  
/mol\_type="genomic DNA"



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US-10-726-967A-3 (1-16) x AR224103 (1-1302)

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 Db 4 ACTCAGCATGTATTGCTCTGCCACTGCCGTACGGGTCTGGGTGTCT 51

RESULT 13 AR269234 1302 bp DNA linear PAT 10-APR-2003  
 LOCUS AR269234  
 DEFINITION Sequence 25 from patent US 6500667.  
 ACCESSION AR269234  
 VERSION AR269234.1 GI:29700202  
 KEYWORDS  
 SOURCE Unknown.  
 ORGANISM Unclassified.

REFERENCE 1 (bases 1 to 1302)  
 AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
 TITLE Aspartyl protease 2 (Asp2) antisense oligonucleotides  
 JOURNAL Patent: US 6500667-A 25 31-DEC-2002;  
 FEATURES Location/Qualifiers  
 source 1..1302  
 /organism="unknown"  
 /mol\_type="genomic DNA"

## ORIGIN

Alignment Scores:  
 Pred. No.: 3.3e-06 Length: 1302  
 Score: 16.00 Matches: 16  
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 Query Match: 100.00% Indels: 0  
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Qy 1 Thrglnhlglylleargleuproleuargserglyleuglyg1yAla 16  
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 Db 4 ACTCAGCATGTATTGCTCTGCCACTGCCGTACGGGTCTGGGTGTCT 51

RESULT 14 AR478789 1302 bp DNA linear PAT 14-MAY-2004  
 LOCUS AR478789  
 DEFINITION Sequence 25 from patent US 6699671.  
 ACCESSION AR478789  
 VERSION AR478789.1 GI:47237509  
 KEYWORDS  
 SOURCE Unknown.  
 ORGANISM Unclassified.

REFERENCE 1 (bases 1 to 1302)  
 AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
 TITLE Alzheimer's disease secretase, APP substrates therefor, and uses therefor  
 JOURNAL Patent: US 6699671-A 25 02-MAR-2004;  
 FEATURES Location/Qualifiers  
 source 1..1302  
 /organism="unknown"  
 /mol\_type="genomic DNA"

## ORIGIN

Alignment Scores:

Pred. No.: 3.3e-06 Length: 1302  
 Score: 16.00 Matches: 16  
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 Query Match: 100.00% Indels: 0  
 DB: 6 Gaps: 0

US-10-726-967A-3 (1-16) x AR478789 (1-1302)

Qy 1 Thrglnhlglylleargleuproleuargserglyleuglyg1yAla 16  
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RESULT 15 AR487355 1302 bp DNA linear PAT 14-MAY-2004  
 LOCUS AR487355  
 DEFINITION Sequence 25 from patent US 6706465.  
 ACCESSION AR487355  
 VERSION AR487355.1 GI:47252453  
 KEYWORDS  
 SOURCE Unknown.  
 ORGANISM Unclassified.

REFERENCE 1 (bases 1 to 1302)  
 AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
 TITLE Method of identifying agents that inhibit APP processing activity  
 JOURNAL Patent: US 6706465-A 25 16-MAR-2004;  
 FEATURES Location/Qualifiers  
 source 1..1302  
 /organism="unknown"  
 /mol\_type="genomic DNA"

## ORIGIN

Alignment Scores:  
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 DB: 6 Gaps: 0

US-10-726-967A-3 (1-16) x AR487355 (1-1302)

Qy 1 Thrglnhlglylleargleuproleuargserglyleuglyg1yAla 16  
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 Db 4 ACTCAGCATGTATTGCTCTGCCACTGCCGTACGGGTCTGGGTGTCT 51

Search completed: July 27, 2005, 19:43:12  
 Job time : 1938 secs

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GenCore version 5.1.6  
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OW protein - nucleic search, using frame\_plus\_p2n model

Run on: July 27, 2005, 17:25:16 ; Search time 432 Seconds  
(without alignments)  
219.250 Million cell updates/sec

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Perfect score: 16  
Sequence: 1 TQHGRLRLRSLGGA 16

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Ygapop 60.0 , Ygapext 60.0  
Fgapop 6.0 , Fgapext 7.0  
Delop 6.0 , Delext 7.0

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Word size: 1

Total number of hits satisfying chosen parameters: 8770599

Minimum DB seq length: 0  
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Prod. No. is the number of results predicted by chance to have a  
score greater than or equal to the score being printed,  
and is derived by analysis of the total score distribution.

# SUMMARIES

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1	16	100.0	1287	4	AAD17895 Human-Asp
2	16	100.0	1287	4	AAD13276 Human-Asp
3	16	100.0	1287	4	AAD06768 Human-Asp
4	16	100.0	1287	4	AA511547 Human-CDN
5	16	100.0	1287	6	AB152487 Human-Asp

6	16	100.0	1287	12	ADJ94362 Human-pro
7	16	100.0	1287	12	AD050458 Human-Asp
8	16	100.0	1287	13	ADR75371 Human-Asp
9	16	100.0	1302	3	AA15670 Human-pro
10	16	100.0	1302	4	AA511713 DNA encod
11	16	100.0	1302	4	AA5117876 Human-pro
12	16	100.0	1302	4	AA51173032 Human-pro
13	16	100.0	1302	4	AA511528 Human-CDN
14	16	100.0	1302	6	AB152468 Human-pro
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16	16	100.0	1302	12	AD050433 Human-pro
17	16	100.0	1302	13	ADR75346 Human-pro
18	16	100.0	1305	4	AA511733 DNA encod
19	16	100.0	1305	4	AA511733 Human-Asp
20	16	100.0	1305	4	AA5117896 Human-Asp
21	16	100.0	1305	4	AA5113277 Human-Asp
22	16	100.0	1305	4	AA511548 Human-CDN
23	16	100.0	1305	4	AA511548 Human-CDN
24	16	100.0	1305	6	AB152488 Human-Asp
25	16	100.0	1305	12	ADJ94364 Human-pro
26	16	100.0	1305	12	AD050460 Human-Asp
27	16	100.0	1305	13	ADR75373 Human-Asp
28	16	100.0	1341	3	AA15668 T7-caspase
29	16	100.0	1341	4	AA511711 DNA encod
30	16	100.0	1341	4	AA5117874 T7-Human-
31	16	100.0	1341	4	AA5113030 T7-Human-
32	16	100.0	1341	4	AA511526 T7-Human-
33	16	100.0	1341	4	AA511526 Human-CDN
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35	16	100.0	1341	12	ADJ94333 Human-CDN
36	16	100.0	1341	12	AD050429 T7-Human-
37	16	100.0	1341	13	ADR75342 T7-Human-
38	16	100.0	1362	3	AA15688 Modified
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40	16	100.0	1362	4	AA5117878 Human-Asp
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42	16	100.0	1362	4	AA511530 Human-Asp
43	16	100.0	1362	4	AA511530 Human-CDN
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RESULT 1	ADJ17895	standard; cDNA; 1287 BP.
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AC	AAAD17895;	
XX	XX	
DT	10-DEC-2001	(first entry)
XX	XX	
DE	Human-Asp 2(b)	protein lacking transmembrane domain encoding cDNA.
XX	XX	
KW	Human; aspartyl protease 2b; Asp2b; amyloid precursor protein; APP;	
KW	Alzheimer's disease; AD; dementia; neurofibrillary tangle; gliosis;	
KW	amyloid plaque; neuronal loss; proteolytic; neurotrophic; neuroprotective;	
KW	ss.	
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OS	Homo sapiens.	
OS	Synthetic.	
XX	XX	
FT	Key	Location/Qualifiers
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FT	FT	/tag= a
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PN	GB2357767-A.	
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PD	04-JUL-2001.	
XX	XX	
PF	22-SEP-2000; 2000GB-00023315.	

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XX 23-SEP-1999; 99US-00404133.
PR 23-SEP-1999; 99US-0155483P.
PR 23-SEP-1999; 99MO-US020881.
PR 13-OCT-1999; 99US-00416901.
PR 06-DEC-1999; 99US-0169232P.
XX
PA (PHAA ) PHARMACIA & UPJOHN CO.
XX
PI Bienkowski MJ, Gurney M;
XX
DR WPI; 2001-444208/48.
P-PSDB; AAE10646.
XX
PT Polypeptide comprising fragments of human aspartyl protease with amyloid
PT precursor protein processing activity and alpha-secretase activity, for
XX identifying modulators useful in treating Alzheimer's disease.
XX
PS Example 10; Page 137; 187p; English.
XX
XX The patent discloses human aspartyl protease 1 (hu-Asp1) or modified Asp1
XX proteins which lack transmembrane domain or amino terminal domain or
XX cytoplasmic domain and retains alpha-secretase activity and amyloid
XX protein precursor (APP) processing activity. The proteins of the
XX invention are useful for assaying hu-Asp1 alpha-secretase activity, which
XX in turn is useful for identifying modulators of hu-Asp1 alpha-secretase
XX activity, where modulators that increase hu-Asp1 alpha-secretase activity
XX are useful for treating Alzheimer's disease (AD) which causes progressive
XX dementia with consequent formation of amyloid plaques, neurofibrillary
XX tangles, gliosis and neuronal loss. Hu-Asp1 protease substrate is useful
XX for assaying hu-Asp1 proteolytic activity, by contacting hu-Asp1 protein
XX with the substrate under acidic conditions and determining the level of
XX hu-Asp1 proteolytic activity. The present sequence is a cDNA encoding
XX human Asp 2(b) protein lacking a transmembrane (TM) domain which is
XX generated by the deletion of the C-terminal TM domain and intracellular
XX domain of human Asp 2(b) protein.
XX
SQ Sequence 1287 BP; 271 A; 370 C; 384 G; 262 T; 0 U; 0 Other;
XX
Alignment Scores:
Pred. No.: 2,64e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
Gaps: 0
DB: 4
XX
US-10-726-967A-3 (1-16) x AAD17895 (1-1287)
OY 1 ThrglnhlsGIYlIeArgLeuProLeuArgSerGlyLeuGIYAla 16
Db 64 ACCGAGCAGCGGCAATCCGGCTCCCTGCGCAGCGGCGGCGGCGCC 111
XX
RESULT 2
ID AAD13276 standard; cDNA; 1287 BP.
XX
AC AAD13276;
XX
DT 23-OCT-2001 (first entry)
XX
DE Human-Asp2(b) deltaTM protein cDNA.
XX
XX Human, aspartyl protease 2b; Asp 2b; beta-amyloid precursor protein; APP;
XX beta-secretase; Alzheimer's disease; dementia; amyloid plaque; gliosis;
XX neurofibrillary tangle; neuronal loss; amyloid-beta peptide; nootropic;
XX neuroprotective; antisense therapy; Asp2(b) deltaTM protein;
XX gene therapy; ss.
XX
OS Homo sapiens.
XX
XX Synthetic.
XX
XX Key Location/Qualifiers

```

```

FT CDS 1..1287
FT /*tag= a
FT /product= "Human Asp2 (b) deltaTM protein"
XX
XX MO200150829-A2.
XX
XX 19-JUL-2001.
XX
XX 09-MAY-2001; 2001MO-IB000799.
XX
XX 09-MAY-2001; 2001MO-IB000799.
XX
XX 09-MAY-2001; 2001MO-IB000799.
XX
XX (BIEN/) BIENKOWSKI M J.
XX (GURN/) GURNEY M E.
XX (HEIN/) HEINRIKSON R L.
XX (PARO/) PARODI L A.
XX (YANR/) YAN R.
XX
XX Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;
XX
XX WPI; 2001-483072/52.
XX
XX P-PSDB; AAE06891.
XX
XX Novel purified polypeptide comprising fragment of mammalian aspartyl
XX protease 2, lacking Asp2 transmembrane domain and retaining beta
XX secretase activity of Asp2 useful for identifying inhibitors of Asp2
XX activity.
XX
XX Example 10; Page 166-167; 185p; English.
XX
XX The invention relates to human aspartyl proteases (Hu-Asp), beta-amyloid
XX precursor protein (APP) isoforms and their corresponding DNA molecules.
XX Human aspartyl proteases can act as beta-secretase proteases useful for
XX treating Alzheimer's disease. APP isoforms are useful for identifying
XX modulators of amyloid-beta peptide production, for use in designing
XX therapeutics for the treatment and prevention of Alzheimer's disease,
XX dementia, formation of amyloid plaques, neurofibrillary tangles, gliosis
XX and neuronal loss. APP isoforms are also used in methods for identifying
XX inhibitors and modulators of human Asp2 activity. The invention relates
XX to a method for identifying agents that modulate the activity of human
XX aspartyl protease Asp2. Amyloid-beta peptides obtained from APP are used
XX as a means to screen in cellular assays for the inhibitors of beta- and
XX gamma- secretase. Hu-Asp DNA fragments are useful as probes or primers in
XX PCR polymerase chain reactions (PCR). The probes are useful for detecting hu-
XX Asp nucleic acids in in vitro assays and in Northern and Southern blots.
XX The present cDNA sequence encodes human aspartyl protease 2b (Hu-Asp2b)
XX deltaTM protein which is obtained by the deletion of C-terminal
XX transmembrane and intracellular domains of Hu-Asp2b. Human Asp2b has beta
XX secretase activity
XX
XX
SQ Sequence 1287 BP; 271 A; 370 C; 384 G; 262 T; 0 U; 0 Other;
XX
Alignment Scores:
Pred. No.: 2,64e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 4 Gaps: 0
XX
US-10-726-967A-3 (1-16) x AAD13276 (1-1287)
OY 1 ThrglnhlsGIYlIeArgLeuProLeuArgSerGlyLeuGIYAla 16
Db 64 ACCGAGCAGCGGCAATCCGGCTCCCTGCGCAGCGGCGGCGGCGCC 111
XX
RESULT 3
ID AAD06768 standard; cDNA; 1287 BP.
XX
AC AAD06768;
XX
DT 10-AUG-2001 (first entry)

```

```
XX Human aspartyl protease 2 (b) delta TM cDNA.
DE
XX
XX Human; alpha-secretase; amyloid precursor protein; APP; therapy;
XX Alzheimer's disease; antialzheimer's; aspartyl protease 2; Asp 2;
XX beta-secretase; chromosome 11q23.3-24.1; mutant; ss.
XX
OS Homo sapiens.
OS Synthetic.
XX
XX Key Location/Qualifiers
XX CDS 1..1287
XX FT /tag= a
XX FT /product= "Human aspartyl protease 2 (b) delta TM"
XX
XX WO200123533-A2.
XX
XX 05-APR-2001.
XX
XX 22-SEP-2000; 2000WO-US026080.
XX
XX 23-SEP-1999; 99US-0155493P.
XX 23-SEP-1999; 99WO-US020881.
XX 13-OCT-1999; 99US-00416901.
XX 06-DEC-1999; 99US-0169232P.
XX
XX (PHAA ) PHARMACIA & UPJOHN CO.
XX
XX Gurney M, Bienkowski MJ;
XX
XX WPI; 2001-290516/30.
XX P-PSDB; AAE02598.
XX
XX Enzymes that cleave the alpha-secretase site of the amyloid precursor
XX protein, useful for the treatment of Alzheimer's disease.
XX
XX Example 10; Page 165-166; 189pp; English.
XX
XX The present invention relates to enzymes for cleaving the alpha-
XX secretase site of the amyloid precursor protein (APP) and methods of
XX identifying those enzymes. The methods may be used to identify enzymes
XX that may be used to cleave the alpha-secretase cleavage site of the APP
XX protein. The enzymes may be used to treat or modulate the progress of
XX Alzheimer's disease. The present sequence is human aspartyl protease 2
XX (Asp 2) (b) delta TM cDNA. The Asp 2 gene from which it is derived is
XX located on chromosome 11q23.3-24.1. The Asp 2 has beta-secretase protease
XX activity
XX
XX Sequence 1287 BP; 271 A; 370 C; 384 G; 262 T; 0 U; 0 Other;
XX
XX Alignment Scores:
XX Pred. No.: 2.64e-06 Length: 1287
XX Score: 16.00 Matches: 16
XX Percent Similarity: 100.00% Conservative: 0
XX Best Local Similarity: 100.00% Mismatches: 0
XX Query Match: 100.00% Indels: 0
XX DB: 4 Gaps: 0
XX
XX US-10-726-967A-3 (1-16) x AAD06768 (1-1287)
XX
XX 1 THRGTHIGGLYLGARGLAEPRLQAVAGSERCYLQNGLYGLYLA 16
XX |||||
XX 64 ACCGACACGCGCATCCGGCTGCCCTGCCAGCGCGCTGGGAGCGCC 111
XX
XX RESULT 4
XX AAS11547
XX ID AAS11547 standard; cDNA; 1287 BP.
XX
XX AAS11547;
XX
XX 24-OCT-2001 (first entry)
XX
XX Human cDNA encoding Human-pro-Asp 2 (b) delta TM.
```

```
XX Human; Aspartyl protease; beta-secretase; nontropic; ASP2;
XX neuroprotective; amyloid protein precursor; APP; Alzheimer's disease;
XX amyloid-beta; Abeta; Human-pro-Asp 2(b) delta TM; ss; mutant.
XX
XX Homo sapiens.
XX Synthetic.
XX
XX Key Location/Qualifiers
XX CDS 1..1287
XX FT /tag= a
XX FT /product= "Human-Pro-Asp 2(b) delta TM"
XX
XX WO200149098-A2.
XX
XX 12-JUL-2001.
XX
XX 09-MAY-2001; 2001WO-IB000798.
XX
XX 09-MAY-2001; 2001WO-IB000798.
XX
XX (BIEN/) BIENKOWSKI M J.
XX (GURN/) GURNEY M E.
XX (HEIN/) HEINRIKSON R L.
XX (PARO/) PARODI L A.
XX (YANR/) YAN R.
XX
XX Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;
XX
XX WPI; 2001-502549/55.
XX
XX Novel purified polypeptide comprising fragment of mammalian aspartyl
XX protease 2, lacking Asp2 transmembrane domain and retaining beta
XX secretase activity of Asp2 useful for identifying inhibitors of Asp2
XX activity.
XX
XX Disclosure; Page 166-167; 185pp; English.
XX
XX The invention relates to a purified polypeptide comprising a fragment of
XX mammalian aspartyl protease (Asp) 2 protein which lacks the Asp2
XX transmembrane domain and the Asp2 protein, and where the polypeptide and
XX the fragment retain the beta-secretase activity of the mammalian Asp2
XX protein. The invention also details polynucleotides for the Asp proteins
XX and vectors expressing them, and a polypeptide (isoform of amyloid
XX protein precursor (APP) comprising the amino acid sequence of an APP or
XX its fragment containing an APP cleavage site recognizable by a mammalian
XX beta-secretase, and further comprising two lysine residues at the
XX carboxyl terminus of the amino acid sequence of the mammalian APP or APP
XX fragment. Also included in the invention are methods of identifying
XX modulators or inhibitors of Asp2. Modulators and inhibitors of Asp2 are
XX useful for treating Alzheimer's disease. APP is useful in methods for
XX identifying inhibitors or modulators of human Asp2 activity and amyloid-
XX beta (Abeta) peptide production. APP is also useful in designing
XX therapeutics for the treatment or prevention of Alzheimer's disease. APP
XX comprising the APP-Sw-beta-secretase peptide sequence (NLDA), which is
XX associated with increased levels of Abeta processing is useful in assays
XX relating the Alzheimer's research. The expression vector is useful for
XX recombinantly expressing APP. Nucleic acids that hybridize to Asp
XX oligonucleotides are useful as probes or primers. The probes are useful
XX for detecting Hu-Asp nucleic acids in in vitro assays and in Northern and
XX Southern blots. The present sequence encodes Human-pro- Asp 2(b) delta TM
XX protein, which lacks the C-terminal transmembrane domain
XX
XX Sequence 1287 BP; 271 A; 370 C; 384 G; 262 T; 0 U; 0 Other;
XX
XX Alignment Scores:
XX Pred. No.: 2.64e-06 Length: 1287
XX Score: 16.00 Matches: 16
XX Percent Similarity: 100.00% Conservative: 0
XX Best Local Similarity: 100.00% Mismatches: 0
XX Query Match: 100.00% Indels: 0
XX DB: 4 Gaps: 0
```

US-10-726-967a-3 (1-16) x AAS11547 (1-1287)

Qy 1 ThglnhlsiglylleArgleuProleuArgSerclyleuGlyyla 16  
Db 64 ACCGAGCAGCGCATCCGCTGCCCCCTGCGAGCGGCTGAGGAGCGCC 111

## RESULT 5

ABL52487  
ID ABL52487 standard; cDNA; 1287 BP.

AC ABL52487;

DT 16-JUL-2002 (first entry)

DE Human Asp-2(b)deltaTM nucleotide sequence SEQ ID NO:50.

KM Human: Asp-1; Asp-2; aspartyl protease; Alzheimer's disease; proteolytic;  
XX chromosome 11q23.3-24.1; gene; ss.

OS Homo sapiens.

XX Key Location/Qualifiers  
FH 1. 1287  
FT CDS /tag= a  
FT /product= "Human Asp-2(b)delta TM"

XX GB2367060-A.

XX 27-MAR-2002.

XX 29-OCT-2001; 2001GB-00025934.

XX 23-SEP-1999; 99US-00404133.

XX 23-SEP-1999; 99US-0155493P.

XX 23-SEP-1999; 99US-0155493P.

XX 13-OCT-1999; 99US-00416901.

XX 06-DEC-1999; 99US-0169232P.

XX 22-SEP-2000; 2000GB-00023315.

XX (PHAA ) PHARMACIA & UPJOHN CO.

XX Bienkowski MJ, Gurney M;

XX WPI: 2002-397167/43.

XX P-PSDB; ABB78607.

Example 10; Page 137; 182pp; English.

The present invention describes a human aspartyl protease 1 (hu-Asp1) substrate (I) which comprises a peptide of no more than 50 amino acids, and which comprises the 8 amino acid sequence Gly-Leu-Ala-Leu-Ala-Leu-Glu-Pro. Also described are: (1) a method (II) for assaying hu-Asp1 proteolytic activity, comprising: (a) contacting a hu-Asp1 protein with (i) under acidic conditions; and (b) determining the level of hu-Asp1 proteolytic activity; (2) a purified polynucleotide (III) comprising a nucleotide sequence that hybridizes under stringent conditions to the non-coding strand complementary to a defined 1804 nucleotide sequence (see ABI52456) where the nucleotide sequence encodes a polypeptide having Asp1 proteolytic activity and lacks nucleotides encoding a transmembrane domain; (3) a purified polynucleotide (III') comprising a sequence that encodes a polypeptide further lacking a pro-peptide domain corresponding to amino acids 23-62 of hu-Asp1 (see ABB78589); (4) a vector (IV) comprising (III) or (III') and (5) a host cell (V) transformed or transfected with (III), (III'), and/or (IV). The hu-Asp1 protease substrate (I) may be used as an enzyme substrate in assays to detect aspartyl protease activity, (II) and therefore diagnose diseases associated with aberrant hu-Asp1 expression and activity such as Alzheimer's disease. Hu-Asp1 has been localized to chromosome 21, while hu-Asp2 has been localized to chromosome 11q23.3-24.1. The present

CC sequence encodes human Asp-2(b)deltaTM, which is given in an example from  
CC the present invention

XX SQ Sequence 1287 BP; 271 A; 370 C; 384 G; 262 T; 0 U; 0 Other;

## Alignment Scores:

Pred. No.:	2,64e-06	Length:	1287
Score:	16.00	Matches:	16
Percent Similarity:	100.00%	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	100.00%	Indels:	0
DB:	6	Gaps:	0

US-10-726-967a-3 (1-16) x ABL52487 (1-1287)

Qy 1 ThglnhlsiglylleArgleuProleuArgSerclyleuGlyyla 16  
Db 64 ACCGAGCAGCGCATCCGCTGCCCCCTGCGAGCGGCTGAGGAGCGCC 111

## RESULT 6

ADJ94362  
ID ADJ94362 standard; cDNA; 1287 BP.

XX ADJ94362;

XX 03-JUN-2004 (first entry)

XX Human-pro-Asp-2(b)deltaTM cDNA.

XX Human; ss; gene; aspartyl protease; Asp-1; Asp-2(a); Asp-2(b);  
XX beta secretase; amyloid protein precursor; APP; Alzheimer's disease;  
XX notropic; neuroprotective; amyloid beta; mutant.

XX Homo sapiens.

XX Synthetic.

XX US6706485-B1.

XX 16-MAR-2004.

XX 12-APR-2000; 2000US-00548376.

XX 24-SEP-1998; 98US-0101594P.

XX 23-SEP-1999; 99US-00404133.

XX 23-SEP-1999; 99US-0155493P.

XX 23-SEP-1999; 99US-0155493P.

XX 13-OCT-1999; 99US-00416901.

XX (PHAA ) PHARMACIA & UPJOHN CO.

XX Gurney ME, Bienkowski MJ, Heinrichson RU, Parodi LA, Yan R;

XX WPI: 2004-236722/22.

XX P-PSDB; ADJ94363.

Identifying agents that modulate activity of Asp2 aspartyl protease  
PT useful for treating or preventing Alzheimer's disease involves comparing  
PT APP processing activity of protease in presence and absence of test  
PT agent.

XX Example 10; SEQ ID NO 50; 109pp; English.

The invention relates to identifying agents that modulate activity of  
CC Asp2 (e.g. a beta-secretase, e.g. human Asp-2(b) appearing as ID 6,  
CC encoded by ID 5) aspartyl protease, involves contacting Asp2 with amyloid  
CC precursor protein (APP) in the presence and absence of a test agent,  
CC where Asp2 is a recombinant polypeptide and processes APP into amyloid  
CC beta, determining APP processing activity of Asp2 in presence and absence  
CC of the test agent, and comparing the activities to identify agents that  
CC modulate the activity of Asp2. Also disclosed are the cDNA and proteins  
CC for human Asp-1 and Asp-2(a) mouse Asp-2(b), a vector comprising the  
CC nucleic acid encoding hu-Asp2 protease sequence, a host cell comprising  
CC the vector and the method of producing hu-Asp polypeptide, an isolated

```
CC antibody that specifically binds to Hu-Ap2 polypeptides, identifying a
CC cell that can be used to screen for inhibitors of beta secretase
CC activity, novel isoforms of amyloid protein precursor (APP), where the
CC last 2 carboxy terminus amino acids of that isoform are both lysine
CC residues (e.g. those designated APP695-KK or carrying the Swedish
CC mutation where KM at 595-596 is mutated to NL, designated e.g. APP695-Sw
CC or APP695-Sw-KK, or a V to F mutation at 642, e.g. APP695-VF, all useful
CC for assaying for beta secretase activity and screening for inhibitors of
CC beta-secretase) and polynucleotides that encode the APP proteins. The
CC method is useful for identifying agents that modulate the activity
CC (amyloid precursor protein processing activity) of Asp2 aspartyl
CC protease. Preferably, the method is useful for identifying agents that
CC inhibit Asp2 aspartyl protease activity. The inhibitors of amyloid
CC precursor protein processing, are useful for treating or preventing
CC Alzheimer's disease. The present sequence encodes an aspartyl protease
CC mutant construct (e.g. lacking a transmembrane domain and/or including a
CC caspase cleavage site) used to investigate the cleavage activity of Asp2
CC proteins.
CC
XX SQ Sequence 1287 BP, 271 A, 370 C, 384 G, 262 T, 0 U, 0 Other;

Alignment Scores:
Pred. No.: 2,64e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 12 Gaps: 0

US-10-726-967A-3 (1-16) x ADJ94362 (1-1287)
QY 1 ThrGlnHisGlyIleArgLeuProLeuArgSerGlyLeuGlyAla 16
DB 64 ACCCAGCAGCGCATCCGCTGCCCTGCGCAGCGCTCGGCGGCC 111

RESULT 7
ADOS0458
ID ADOS0458 standard; DNA; 1287 BP.
XX
AC ADOS0458;
XX
DT 29-JUL-2004 (first entry)
XX
DE Human Asp2(b)deltaTM mutant DNA.
XX
KM Aspartyl protease; Asp; beta secretase; amyloid precursor protein; APP;
XX Alzheimer's disease; gene therapy; human; mutant; gene; de.
XX
OS Homo sapiens.
OS Synthetic.
XX
FH Key Location/Qualifiers
FT CDS 1..1287
FT /tag= a
FT /product= "Human Asp2(b) mutant protein"
XX
XX US6737510-B1.
XX
XX PD 18-MAY-2004.
XX
XX PF 12-APR-2000; 2000US-00548373.
XX
XX 24-SEP-1998; 98US-0101594P.
XX 23-SEP-1999; 99US-00404133.
XX 23-SEP-1999; 99US-0155493P.
XX 23-SEP-1999; 99MO-US020881.
XX 13-OCT-1999; 99US-00416901.
XX
XX (PHAA ) PHARMACIA & UPJOHN CO.
XX
XX Gurney ME, Bienkowski MJ, Heinrikson RL, Parodi LA, Yan R;
XX
XX WPI; 2004-387112/36.
```

```
DR P-PSDB; ADOS0459.
XX
XX New Asp2 aspartyl protease protein comprising tripeptides DTG and DSG
PT involved in processing amyloid precursor protein into amyloid beta,
PT useful in preparing a composition for treating or preventing Alzheimer's
PT disease.
XX
XX Example 10; SEQ ID NO 50; 108bp; English.
XX
XX The invention relates to a method for identifying an agent that decreases
CC the protease activity of the aspartyl protease (APP) polypeptide. It also
CC provides enzyme and enzymatic procedures for cleaving the beta secretase
CC cleavage site of the amyloid precursor protein (APP). The invention is
CC useful in preparing a composition for treating or preventing Alzheimer's
CC disease. It is also useful in gene therapy. The present sequence is human
CC Asp2(b) mutant DNA. This sequence is used to illustrate the method of the
CC invention.
CC
XX SQ Sequence 1287 BP, 271 A, 370 C, 384 G, 262 T, 0 U, 0 Other;

Alignment Scores:
Pred. No.: 2,64e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 12 Gaps: 0

US-10-726-967A-3 (1-16) x ADOS0458 (1-1287)
QY 1 ThrGlnHisGlyIleArgLeuProLeuArgSerGlyLeuGlyAla 16
DB 64 ACCCAGCAGCGCATCCGCTGCCCTGCGCAGCGCTCGGCGGCC 111

RESULT 8
ADR75371
ID ADR75371 standard; DNA; 1287 BP.
XX
AC ADR75371;
XX
DT 18-NOV-2004 (first entry)
XX
DE Human Asp2(b)deltaTM mutant DNA.
XX
KM Aspartyl protease; Asp; amyloid precursor protein; APP; amyloid beta;
XX chromosome identification; Alzheimer's disease; human; mutant; gene; de.
XX
OS Homo sapiens.
OS Synthetic.
XX
FH Key Location/Qualifiers
FT CDS 1..1287
FT /tag= a
FT /product= "Human Asp2(b) mutant protein"
XX
XX US2004166507-A1.
XX
XX PD 26-AUG-2004.
XX
XX PF 29-AUG-2003; 2003US-00652045.
XX
XX 24-SEP-1998; 98US-0101594P.
XX 23-SEP-1999; 99US-00404133.
XX 23-SEP-1999; 99US-0155493P.
XX 13-OCT-1999; 99US-00416901.
XX
XX (GURN/) GURNEY M E.
XX (BIEN/) BIENKOWSKI M J.
XX (HEIN/) HEINRIKSON R L.
XX (PARO/) PARODI L A.
XX (YANR/) YAN R.
XX
XX Gurney ME, Bienkowski MJ, Heinrikson RL, Parodi LA, Yan R;
XX
XX
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```
XX WPI; 2004-624916/60.
DR P-PSDB; ADR75372.
XX
PT Novel purified/isolated polynucleotide encoding polypeptide having
PT aspartyl protease activity involved in processing amyloid precursor
PT protein into amyloid beta, useful in identifying agent decreasing
PT activity of aspartyl protease.
XX
XX Example 10; SEQ ID NO 50; 107bp; English.
XX
XX The invention relates to nucleic acid sequences encoding aspartyl
XX protease (Asp) polypeptides having aspartyl protease activity involved in
XX processing amyloid precursor protein (APP) into amyloid beta. The
XX invention also relates to a method for identifying an agent that
XX decreases the protease activity of the Asp. Asp DNA is useful in
XX chromosome identification as they can hybridize with a specific location
XX on a human chromosome and in identifying the relationship between genes
XX and diseases (particular gene responsible for causing diseases). It is
XX also useful for identifying candidates to modulate the progression of
XX Alzheimer's disease. Asp is useful in raising antibodies that are useful
XX in diagnostic assay for detecting Hu-Asp polypeptide expression. The
XX present sequence is the human Asp2(b) deltaTM mutant DNA. This sequence is
XX used to illustrate the method of the invention.
XX
SQ Sequence 1287 BP; 271 A; 370 C; 384 G; 262 T; 0 U; 0 Other;
XX
Alignment Scores:
Pred. No.: 2.64e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 13 Gaps: 0
XX
US-10-726-967A-3 (1-16) x ADR75371 (1-1287)
XX
OY 1 ThGlnHIGlyYlLeArgLeuProLeuArgSerGlyLeuGlyGlyAla 16
Db 64 ACCACACACGCGATCCGCTGCTGCCCTGCGACGCGCTGGGGGCGCC 111
XX
RESULT 9
ID AA15670 standard; DNA; 1302 BP.
XX
AC AA15670;
XX
DT 15-SEP-2003 (revised)
DT 06-AUG-2003 (revised)
DT 03-AUG-2000 (first entry)
XX
DE Human-pro-Asp-2(a)-deltaTM nucleotide sequence.
XX
KW Aspartyl protease; aspartase; amyloid precursor protein; APP; Asp 2; ss;
KW Alzheimer's disease; beta secretase site; human-pro-Asp-2(a)-deltaTM.
XX
OS Homo sapiens.
OS Enterobacteria phage T7.
XX
XX WO200017369-A2.
XX
XX 30-MAR-2000.
XX
XX 23-SEP-1999; 99WO-US020881.
XX
XX 24-SEP-1998; 98US-0101594P.
XX
XX (PhA ) PHARMACIA & UPJOHN CO.
XX
XX Gurney ME, Bienkowski MJ, Heintikson RL, Parodi LA, Yan R;
XX WPI; 2000-303209/26.
XX
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```
DR P-PSDB; AAY98433.
XX
XX New enzyme designated human aspartase useful in research into Alzheimer's
XX disease is capable of cleaving amyloid protein precursor at the beta
XX secretase site to produce amyloid beta peptide.
XX
XX Example 9; Fig 8; 183bp; English.
XX
XX This sequence represents a modified version of the human aspartase 2
XX (Asp2) nucleotide sequence. The sequence is used in the bacterial
XX expression of human Asp2L. The invention relates to a protease (e.g.,
XX Asp2) capable of cleaving the beta secretase site of amyloid precursor
XX protein (APP). The protease contains a sequence encoding the amino acid
XX sequence DNG and a sequence encoding DSG or DNG separated by 100-300
XX amino acids. When mutated the Asp gene causes an autosomal dominant form
XX of Alzheimer's disease. APP localises to the cell surface membrane and
XX have a single C-terminal transmembrane domain. Proteolytic processing of
XX APP produces the amyloid beta protein, which is possibly very important
XX in Alzheimer's disease. The invention includes a nucleotide sequence
XX encoding the protease, a vector containing the nucleotide sequence, and a
XX cell line comprising the vector. Methods for screening for inhibitors of
XX aspartase protein and nucleotide sequences and the methods for
XX identifying inhibitors of the protease, are useful in the treatment of
XX and research in to Alzheimer's disease. (Updated on 06-AUG-2003 to
XX correct OS field.) (Updated on 15-SEP-2003 to standardise OS field)
XX
SQ Sequence 1302 BP; 281 A; 367 C; 370 G; 284 T; 0 U; 0 Other;
XX
Alignment Scores:
Pred. No.: 2.67e-06 Length: 1302
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 3 Gaps: 0
XX
US-10-726-967A-3 (1-16) x AA15670 (1-1302)
XX
OY 1 ThGlnHIGlyYlLeArgLeuProLeuArgSerGlyLeuGlyGlyAla 16
Db 4 ACTCAGCATGCTATTGCTGTCGCACTGCGTAGCGGTCTGGGTGGTGGCT 51
XX
RESULT 10
ID AAS11713
XX
AC AAS11713;
XX
DT 11-SEP-2003 (revised)
DT 24-OCT-2001 (first entry)
XX
DE DNA encoding T7-human aspartyl protease 2a deltaTM (10w GC).
XX
KW Human; aspartyl protease 1; Asp-1; nootropic; neuroprotective;
KW aspartyl protease 2; Asp2; amyloid protein precursor; APP;
KW beta-secretase; Alzheimer's disease; de.
XX
XX Homo sapiens.
XX
XX Enterobacteria phage T7.
XX
XX Key Location/Qualifiers
XX CDS 1..1302
XX FT /*tag= a
XX /product= "T7-Aspartyl protease-2a delta TM (10w GC)"
XX
XX 12-JUL-2001.
XX
XX 09-MAY-2001; 2001WO-IB000797.
XX
XX 09-MAY-2001; 2001WO-IB000797.
XX
```



```
XX (BIEN/) BIENKOWSKI M J.
PA (GURNEY) GURNEY M B.
PA (HEIN/) HEINRIKSON R L.
PA (PARO/) PARODI L A.
PA (YANR/) YAN R.
PI Bienkowski MJ, Gurney ME, Heinrichson RU, Parodi LA, Yan R;
DR WPI; 2001-502548/55.
DR P-PSDB; AAU07113.
XX
XX Novel purified polypeptide comprising fragment of mammalian aspartyl
PT protease 2, lacking Asp2 transmembrane domain and retaining beta
PT secretase activity of Asp2 useful for identifying inhibitors of Asp2
PT activity.
XX
XX Example 9; Fig 8; 185pp; English.
XX
XX The invention relates to a novel purified polypeptide comprising a
CC fragment of mammalian aspartyl protease 2 (Asp2) protein which lacks the
CC Asp2 transmembrane domain and the Asp2 protein, and where the polypeptide
CC and the fragment retain the beta-secretase activity of the mammalian Asp2
CC protein. Also included is an isoform of amyloid protein precursor (APP)
CC comprising the amino acid sequence of a APP or its fragment containing an
CC APP cleavage site recognisable by a mammalian beta-secretase, and further
CC comprising two lysine residues at the carboxyl terminus of the amino acid
CC sequence of the mammalian APP or APP fragment. The polypeptides are used
CC for assaying for modulators of beta-secretase activity; identifying
CC agents that inhibit the APP processing activity of human Asp2 aspartyl
CC protease (Hu-Asp2); identifying agents that modulate the activity of Asp2
CC ; and for reducing cellular production of amyloid beta (Abeta) from APP.
CC Agents identified by the above methods are useful for treating
CC Alzheimer's disease; for identifying modulators of amyloid-beta (Abeta)
CC peptide production, and for use in designing therapeutics for the
CC treatment or prevention of Alzheimer's disease. Probes and primers
CC derived from Asp nucleic acid sequences are useful for detecting Hu-Asp
CC nucleic acids in in vitro assays and in Northern and Southern blots. The
CC present sequence represents the coding sequence of T7-human Asp2a delta
CC TM (low GC) construct which has a T7 tag, has the GC content of the 5'
CC sequence reduced by site-directed mutagenesis, and lacks the
CC transmembrane domain. This construct was used for bacterial expression
CC and purification of human Asp2a. (Updated on 11-SEP-2003 to standardise
CC OS field)
XX
XX SQ Sequence 1302 BP; 281 A; 367 C; 370 G; 284 T; 0 U; 0 Other;
XX
XX Alignment Scores:
XX Pred. No.: 2.67e-06 Length: 1302
XX Score: 16.00 Matches: 16
XX Percent Similarity: 100.00% Conservative: 0
XX Best Local Similarity: 100.00% Mismatches: 0
XX Query Match: 100.00% Indels: 0
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XX US-10-726-967A-3 (1-16) x AAS11713 (1-1302)
XX
XX QY 1 ThrGlnHieGlyTLeaRgLeuProLeuAArgSerGlyLeuGlyVala 16
XX |||||
XX Db 4 ACTCAGCATGATGTCGTCTGCCACTGCGTACGGGTCTGGGTGCTCT 51
XX
XX RESULT 11
XX AAD17876
XX AAD17876 standard; cDNA; 1302 BP.
XX
XX AAD17876;
XX
XX 10-DEC-2001 (first entry)
XX
XX Human-pro-Asp 2(a) protein lacking TM domain (low GC) encoding cDNA.
XX
XX Human: aspartyl protease 1; Asp1; amyloid precursor protein; APP;
XX Alzheimer's disease; AD; dementia; neurofibrillary tangle; gliosis;
XX
```

```
KW amyloid plaque; neuronal loss; proteolytic; nootropic; neuroprotective;
KW Human-pro-Asp 2(a) protein; ss.
XX
XX Homo sapiens.
XX Synthetic.
XX
XX Key Location/Qualifiers
XX CDS 1..1302
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XX FT /product= "Human-pro-Asp 2(a) protein lacking
XX FT transmembrane domain"
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XX GB2357767-A.
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XX 04-JUL-2001.
XX
XX 22-SEP-2000; 2000GB-00023315.
XX
XX 23-SEP-1999; 99US-00404133.
XX 23-SEP-1999; 99US-0155493P.
XX 23-SEP-1999; 99WO-US020881.
XX 13-OCT-1999; 99US-00416901.
XX 06-DEC-1999; 99US-0169232P.
XX
XX (PHAA ) PHARMACIA & UPJOHN CO.
XX
XX PI Bienkowski MJ, Gurney M;
XX
XX WPI; 2001-444208/48.
XX
XX P-PSDB; AAE10640.
XX
XX Polypeptide comprising fragments of human aspartyl protease with amyloid
PT precursor protein processing activity and alpha-secretase activity, for
PT identifying modulators useful in treating Alzheimer's disease.
XX
XX Example 9; Fig 8; 187pp; English.
XX
XX The patent discloses human aspartyl protease 1 (hu-Asp1) or modified Asp1
CC proteins which lack transmembrane domain or amino terminal domain or
CC cytoplasmic domain and retains alpha-secretase activity and amyloid
CC protein precursor (APP) processing activity. The proteins of the
CC invention are useful for assaying hu-Asp1 alpha-secretase activity, which
CC in turn is useful for identifying modulators of hu-Asp1 alpha-secretase
CC activity, where modulators that increase hu-Asp1 alpha-secretase activity
CC are useful for treating Alzheimer's disease (AD) which causes progressive
CC dementia with consequent formation of amyloid plaques, neurofibrillary
CC tangles, gliosis and neuronal loss. Hu-Asp1 protease substrate is useful
CC for assaying hu-Asp1 proteolytic activity, by contacting hu-Asp1 protein
CC with the substrate under acidic conditions and determining the level of
CC hu-Asp1 proteolytic activity. The present sequence is a cDNA encoding
CC human-pro-Asp 2(a) protein lacking a transmembrane (TM) domain (low GC)
CC which is generated from human Asp 2(a) protein by the deletion of its C-
CC terminal transmembrane domain and change of degenerate codons bases in 15
XX amino acid positions from G/C to A/T to reduce the GC content
XX
XX SQ Sequence 1302 BP; 281 A; 367 C; 370 G; 284 T; 0 U; 0 Other;
XX
XX Alignment Scores:
XX Pred. No.: 2.67e-06 Length: 1302
XX Score: 16.00 Matches: 16
XX Percent Similarity: 100.00% Conservative: 0
XX Best Local Similarity: 100.00% Mismatches: 0
XX Query Match: 100.00% Indels: 0
XX DB: 4 Gaps: 0
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XX US-10-726-967A-3 (1-16) x AAD17876 (1-1302)
XX
XX QY 1 ThrGlnHieGlyTLeaRgLeuProLeuAArgSerGlyLeuGlyVala 16
XX |||||
XX Db 4 ACTCAGCATGATGTCGTCTGCCACTGCGTACGGGTCTGGGTGCTCT 51
XX
XX RESULT 12
XX AAD13032
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ID AAD13032 standard; cDNA; 1302 BP.  
 AC AAD13032;  
 DT 23-OCT-2001 (first entry)  
 XX Human-pro-Asp2(a) deltaTM (low GC) protein cDNA.  
 DE Human; aspartyl protease 2a; Asp 2a; beta-amyloid precursor protein; APP;  
 KM beta-secretase; Alzheimer's disease; dementia; amyloid plaque; gliosis;  
 KM neurofibrillary tangle; neuronal loss; amyloid-beta peptide; nototropic;  
 KM neuroprotective; antisense therapy; pro-Asp2(a) deltaTM protein;  
 KM gene therapy; ss.  
 XX Homo sapiens.  
 OS Synthetic.  
 FH Key Location/Qualifiers  
 FT CDS 1..1302  
 FT /\*tag= a  
 FT /product= "Human-pro-Asp2(a) deltaTM (low GC) protein"  
 XX WO200150829-A2.  
 XX 19-JUL-2001.  
 XX 09-MAY-2001; 2001WO-IB000799.  
 XX 09-MAY-2001; 2001WO-IB000799.  
 XX (BIEN/) BIENKOWSKI M J.  
 XX (GUEN/) GUENEY M E.  
 XX (HEIN/) HEINRIKSON R L.  
 XX (PARO/) PARODI L A.  
 XX (YANR/) YAN R.  
 PI Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;  
 XX WPI; 2001-483072/52.  
 DR P-PSDB; AAE06870.  
 XX Novel purified polypeptide comprising fragment of mammalian aspartyl  
 PT proasease 2, lacking Asp2 transmembrane domain and retaining beta  
 PT secretase activity of Asp2 useful for identifying inhibitors of Asp2  
 PT activity.  
 XX Example 9; Fig 8; 185pp; English.  
 XX The invention relates to human aspartyl proteases (Hu-Asp), beta-amyloid  
 CC precursor protein (APP) isoforms and their corresponding DNA molecules.  
 CC Human aspartyl proteases can act as beta-secretase proteases useful for  
 CC treating Alzheimer's disease. APP isoforms are useful for identifying  
 CC modulators of amyloid-beta peptide production, for use in designing  
 CC therapeutics for the treatment and prevention of Alzheimer's disease,  
 CC dementia, formation of amyloid plaques, neurofibrillary tangles, gliosis  
 CC and neuronal loss. APP isoforms are also used in methods for identifying  
 CC inhibitors and modulators of human Asp2 activity. The invention relates  
 CC to a method for identifying agents that modulate the activity of human  
 CC aspartyl protease Asp2. Amyloid-beta peptides obtained from APP are used  
 CC as a means to screen in cellular assays for the inhibitors of beta- and  
 CC gamma- secretase. Hu-Asp DNA fragments are useful as probes or primers in  
 CC polymerase chain reactions (PCR). The probes are useful for detecting Hu-  
 CC Asp nucleic acids in *in vitro* assays and in Northern and Southern blots.  
 CC The present cDNA sequence encodes human-pro-aspartyl protease 2a (Asp2a)  
 CC deltaTM (low GC) protein which is obtained by the deletion of C-terminal  
 CC transmembrane domain and change of degenerate codons bases in 15 amino  
 CC acid positions from G/C to A/T in the Hu-Asp2a. Human Asp2a has beta-  
 CC secretase activity  
 XX Sequence 1302 BP; 281 A; 367 C; 370 G; 284 T; 0 U; 0 Other;  
 SQ Alignment Scores: 2.67e-06 Length: 1302  
 Pred. No.:

Score: 16.00 Matches: 16  
 Percent Similarity: 100.00% Conservative: 0  
 Best Local Similarity: 100.00% Mismatches: 0  
 Query Match: 100.00% Indels: 0  
 DB: 4 Gaps: 0  
 US-10-726-967A-3 (1-16) x AAD13032 (1-1302)  
 QY 1 ThGlnHtAgLy1leaNgLeuPProLeuArgSerGlyLeuGlyAla 16  
 DB 4 ACTCAGATGATGATTCGTCTGCCACCTGCGTAGCGGCTGTGGATGCT 51  
 RESULT 13  
 ID AAD06750 standard; cDNA; 1302 BP.  
 AC AAD06750;  
 DT 10-AUG-2001 (first entry)  
 XX Human-pro-Asp-2(a) deltaTM protein cDNA.  
 DE Human; alpha-secretase; amyloid precursor protein; APP; therapy;  
 KM Alzheimer's disease; anti-Alzheimer's; aspartyl protease 2a; Asp2a;  
 KM beta-secretase; Asp-2a delta TM; ss.  
 XX Homo sapiens.  
 OS Synthetic.  
 FH Key Location/Qualifiers  
 FT CDS 1..1302  
 FT /\*tag= a  
 FT /product= "Human-pro-Asp-2(a) delta TM protein"  
 XX WO200123533-A2.  
 XX 05-APR-2001.  
 XX 22-SEP-2000; 2000WO-US026080.  
 XX 23-SEP-1999; 99US-0155493P.  
 XX 23-SEP-1999; 99WO-US020881.  
 XX 13-OCT-1999; 99US-00416901.  
 XX 06-DEC-1999; 99US-0169232P.  
 XX (PHAA ) PHARMACIA & UPJOHN CO.  
 PI Gurney M, Bienkowski MJ;  
 XX WPI; 2001-290516/30.  
 DR P-PSDB; AAE02592.  
 XX Enzymes that cleave the alpha-secretase site of the amyloid precursor  
 PT protein, useful for the treatment of Alzheimer's disease.  
 XX Example 9; Page 155; 189pp; English.  
 XX The present invention relates to enzymes for cleaving the alpha-  
 CC secretase site of the amyloid precursor protein (APP) and methods of  
 CC identifying those enzymes. The methods may be used to identify enzymes  
 CC that may be used to cleave the alpha-secretase cleavage site of the APP  
 CC protein. The enzymes may be used to treat or modulate the progress of  
 CC Alzheimer's disease. The present sequence is a cDNA encoding human  
 CC aspartyl protease 2a (Asp-2a) deltaTM protein which is obtained by  
 CC deleting the transmembrane domain and adding a T7 tag at the N-terminal  
 CC end. This sequence has beta-secretase protease activity. Note: The  
 CC present sequence is also shown in figure 8 of the specification, but  
 CC lacks nucleotides at its 3' end. This sequence shown in figure 8 has a  
 CC stop codon at its 3' end  
 XX Sequence 1302 BP; 281 A; 367 C; 370 G; 284 T; 0 U; 0 Other;  
 SQ Alignment Scores:

Pred. No.: 2.67e-06 Length: 1302  
 Score: 16.00 Matches: 16  
 Percent Similarity: 100.00% Conservative: 0  
 Best Local Similarity: 100.00% Mismatches: 0  
 Query Match: 100.00% Indels: 0  
 DB: 4 Gaps: 0

US-10-726-967A-3 (1-16) x AAD06750 (1-1302)

Qy 1 ThrGlnHisGlyIleArgLeuProLeuArgSerGlyLeuGlyAla 16  
 Db 4 ACTCAGCATGATTCGTCTGCCACTGCTAGCGGTCTGGTGTCT 51

RESULT 14  
 AAS11528 standard; cDNA; 1302 BP.  
 XX AAS11528;  
 AC AAS11528;  
 XX  
 DT 24-OCT-2001 (first entry)  
 XX  
 DE Human cDNA encoding Human-pro-Asp 2(a) delta TM (low GC).  
 XX  
 KW Human; Aspartyl protease; beta-secretase; neurotropic; ASP2;  
 KM neuroprotective; amyloid protein precursor; APP; Alzheimer's disease;  
 KM amyloid-beta; Abeta; ss; Human-pro-Asp 2(a) delta TM (low GC).  
 XX  
 OS Homo sapiens.  
 OS Synthetic.  
 OS  
 FH Key Location/Qualifiers  
 FT CDS 1..1302  
 FT /tag= a  
 FT /product= "Human-pro-Asp 2(a) delta TM (low GC)"  
 XX  
 PN WO200149098-A2.  
 XX  
 PD 12-JUL-2001.  
 XX  
 PF 09-MAY-2001; 2001WO-IB000798.  
 XX  
 PR 09-MAY-2001; 2001WO-IB000798.  
 XX  
 PA (BIEN/) BIERKOWSKI M J.  
 PA (GURN/) GURNEY M E.  
 PA (HEIN/) HEINRIKSON R L.  
 PA (PARO/) PARODI L A.  
 PA (YANR/) YAN R.  
 XX  
 PI Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;  
 XX  
 DR MPI; 2001-502549/55.  
 XX  
 DR P-PSDB; AAU06614.  
 XX  
 PT Novel purified polypeptide comprising fragment of mammalian aspartyl  
 PT protease 2, lacking Asp2 transmembrane domain and retaining beta  
 PT secretase activity of Asp2 useful for identifying inhibitors of Asp2  
 PT activity.  
 XX  
 PS Example 9; Fig 8; 185pp; English.  
 XX  
 CC The invention relates to a purified polypeptide comprising a fragment of  
 CC mammalian aspartyl protease (Asp)2 protein which lacks the Asp2  
 CC transmembrane domain and the Asp2 protein, and where the polypeptide and  
 CC the fragment retain the beta-secretase activity of the mammalian Asp2  
 CC protein. The invention also details polynucleotides for the Asp protease  
 CC and vectors expressing them, and a polypeptide isoform of amyloid  
 CC protein precursor (APP) comprising the amino acid sequence of an APP or  
 CC its fragment containing an APP cleavage site recognizable by a mammalian  
 CC beta-secretase, and further comprising two lysine residues at the  
 CC carboxyl terminus of the amino acid sequence of the mammalian APP or APP  
 CC fragment. Also included in the invention are methods of identifying  
 CC modulators or inhibitors of Asp2. Modulators and inhibitors of Asp2 are

CC useful for treating Alzheimer's disease. APP is useful in methods for  
 CC identifying inhibitors or modulators of human Asp2 activity and amyloid-  
 CC beta (Abeta) peptide production. APP is also useful in designing  
 CC therapeutics for the treatment or prevention of Alzheimer's disease. APP  
 CC comprising the APP-Sw-beta-secretase peptide sequence (NDA), which is  
 CC associated with increased levels of Abeta processing is useful in assays  
 CC relating the Alzheimer's research. The expression vector is useful for  
 CC recombinantly expressing APP. Nucleic acids that hybridize to Asp  
 CC oligonucleotides are useful as probes or primers. The probes are useful  
 CC for detecting Hu-Asp nucleic acids in in vitro assays and in Northern and  
 CC Southern blots. The present sequence encodes Human-pro-Asp 2(a) delta TM  
 CC (low GC), a synthetic version of Asp 2(a) whose GC content has been  
 CC altered to facilitate expression in E.coli

XX Sequence 1302 BP; 281 A; 367 C; 370 G; 284 T; 0 U; 0 Other;

Alignment Scores:  
 Pred. No.: 2.67e-06 Length: 1302  
 Score: 16.00 Matches: 16  
 Percent Similarity: 100.00% Conservative: 0  
 Best Local Similarity: 100.00% Mismatches: 0  
 Query Match: 100.00% Indels: 0  
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US-10-726-967A-3 (1-16) x AAS11528 (1-1302)

Qy 1 ThrGlnHisGlyIleArgLeuProLeuArgSerGlyLeuGlyAla 16  
 Db 4 ACTCAGCATGATTCGTCTGCCACTGCTAGCGGTCTGGTGTCT 51

RESULT 15  
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 ID ABL52468 standard; cDNA; 1302 BP.  
 XX ABL52468;  
 AC ABL52468;  
 XX  
 DT 16-JUL-2002 (first entry)  
 XX  
 DE Human-pro-Asp-2(a)deltaTM (low GC) nucleotide sequence SEQ ID NO:25.  
 XX  
 KW Human; Asp-1; Asp-2; aspartyl protease; Alzheimer's disease; proteolytic;  
 KM amyloid precursor protein; APP; gene; ss.  
 XX  
 OS Homo sapiens.  
 OS  
 FH Key Location/Qualifiers  
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 FT /tag= a  
 FT /product= "human-pro-Asp-2(a)deltaTM (low GC)"  
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 PN GB2367060-A.  
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 PD 27-MAR-2002.  
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 PF 29-OCT-2001; 2001GB-00025934.  
 XX  
 PR 23-SEP-1999; 99US-00404133.  
 PR 23-SEP-1999; 99US-0155493P.  
 PR 23-SEP-1999; 99WO-US020881.  
 PR 13-OCT-1999; 99US-00416901.  
 PR 06-DEC-1999; 99US-0169232P.  
 PR 22-SEP-2000; 2000GB-00023315.  
 XX  
 PA (PHAA ) PHARMACIA & UPJOHN CO.  
 XX  
 PI Bienkowski MJ, Gurney M;  
 XX  
 DR MPI; 2002-397167/43.  
 DR P-PSDB; ABB78601.  
 XX  
 PT Human aspartyl protease 1 substrates useful in assays to detect aspartyl  
 PT protease activity, e.g. for the diagnosis of Alzheimer's disease.  
 XX

PS Example 9; Fig 8; 182pp; English.

XX  
 CC The present invention describes a human aspartyl protease 1 (hu-Asp1)  
 CC substrate (I) which comprises a peptide of no more than 50 amino acids,  
 CC and which comprises the 8 amino acid sequence Gly-Leu-Ala-Leu-Ala-Leu-  
 CC Glu-Pro. Also described are: (1) a method (II) for assaying hu-Asp1  
 CC proteolytic activity, comprising: (a) contacting a hu-Asp1 protein with  
 CC (1) under acidic conditions; and (b) determining the level of hu-Asp1  
 CC proteolytic activity; (2) a purified polynucleotide (III) comprising a  
 CC nucleotide sequence that hybridises under stringent conditions to the non  
 CC -coding strand complementary to a defined 1804 nucleotide sequence (see  
 CC ABUS2456) where the nucleotide sequence encodes a polypeptide having Asp1  
 CC proteolytic activity and lacks nucleotides encoding a transmembrane  
 CC domain; (3) a purified polynucleotide (III') comprising a sequence that  
 CC hybridises under stringent conditions to (III) (the nucleotide sequence  
 CC encodes a polypeptide further lacking a pro-peptide domain corresponding  
 CC to amino acids 23-62 of hu-Asp1 (see ABUS2456)); (4) a vector (IV)  
 CC comprising (III) or (III'); and (5) a host cell (V) transformed or  
 CC transfected with (III), (III') and/or (IV). The hu-Asp1 protease  
 CC substrate (1) may be used as an enzyme substrate in assays to detect  
 CC aspartyl protease activity, (II) and therefore diagnose diseases  
 CC associated with aberrant hu-Asp1 expression and activity such as  
 CC Alzheimer's disease. Hu-Asp1 has been localised to chromosome 21, while  
 CC hu-Asp2 has been localised to chromosome 11q23.3-24.1. The present  
 CC sequence encodes human-pro-Asp-2(a)delatm (low GC), which is given in an  
 CC example from the present invention  
 XX

SQ Sequence 1302 BP; 281 A; 367 C; 370 G; 284 T; 0 U; 0 Other;

Alignment Scores:

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US-10-726-967A-3 (1-16) x ABUS2468 (1-1302)

QY	1	ThRGInHISGLYIleargleuProleuAargSergIyleuGIYGIYAla	16
DB	4	ACTCAGCATGGTATTCGTCTGCCTGCGTAGCGGTCTGGGTGGTCT	51

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GenCore version 5.1.6  
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Title: US-10-726-967A-3

Perfect score: 16  
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Total number of hits satisfying chosen parameters: 2397162

Minimum DB seq length: 0

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score greater than or equal to the score of the result being printed,  
and is derived by analysis of the total score distribution.

## SUMMARIES

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5	16	100.0	1287	4	US-09-416-901B-50
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7	16	100.0	1287	4	US-09-794-927A-50
8	16	100.0	1287	4	US-09-548-373D-50
9	16	100.0	1287	4	US-09-869-414-50
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24	16	100.0	1302	4	US-09-806-194A-25	Sequence 25, Appl
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31	16	100.0	1305	4	US-09-794-927A-52	Sequence 52, Appl
32	16	100.0	1305	4	US-09-548-373D-52	Sequence 52, Appl
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34	16	100.0	1305	4	US-09-869-414-52	Sequence 52, Appl
35	16	100.0	1305	4	US-09-548-366F-52	Sequence 52, Appl
36	16	100.0	1305	4	US-09-548-368D-52	Sequence 52, Appl
37	16	100.0	1305	4	US-09-794-925A-52	Sequence 52, Appl
38	16	100.0	1341	3	US-09-548-372D-21	Sequence 21, Appl
39	16	100.0	1341	3	US-09-548-372D-21	Sequence 21, Appl
40	16	100.0	1341	4	US-09-551-853D-21	Sequence 21, Appl
41	16	100.0	1341	4	US-09-416-901B-21	Sequence 21, Appl
42	16	100.0	1341	4	US-09-548-376D-21	Sequence 21, Appl
43	16	100.0	1341	4	US-09-794-927A-21	Sequence 21, Appl
44	16	100.0	1341	4	US-09-548-373D-21	Sequence 21, Appl
45	16	100.0	1341	4	US-09-795-847B-21	Sequence 21, Appl

## ALIGNMENTS

RESULT 1  
US-09-548-372D-50  
; Sequence 50, Application US/09548372D  
; Patent No. 6420534  
; GENERAL INFORMATION:  
; APPLICANT: GUNNEY ET AL.  
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES  
; TITLE OF INVENTION: THEREOF  
; FILE REFERENCE: 29915/62801  
; CURRENT APPLICATION NUMBER: US/09/548,372D  
; PRIOR FILING DATE: 2000-04-12  
; PRIOR APPLICATION NUMBER: US 60/155,493  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: US 09/404,133  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: PCT/US99/20881  
; PRIOR FILING DATE: 1999-09-23  
; PRIOR APPLICATION NUMBER: US 60/101,594  
; PRIOR FILING DATE: 1998-09-24  
; NUMBER OF SEQ ID NOS: 73  
; SOFTWARE: PatentIn version 3.1  
; SEQ ID NO 50  
; LENGTH: 1287  
; TYPE: DNA  
; ORGANISM: Artificial sequence  
; FEATURES:  
; OTHER INFORMATION: Hu-Asp2(b) delta TM  
US-09-548-372D-50  
Alignment Scores:  
Pred. No.: 1.48e-06  
Score: 16.00  
Percent Similarity: 100.00%  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
DB: 3  
Length: 1287  
Matches: 16  
Conservative: 0  
Mismatch: 0  
Indels: 0  
Gaps: 0

US-10-726-967A-3 (1-16) x US-09-548-372D-50 (1-1287)

QY 1 ThrglnhsglylIleArgLeuProLeuArgSerGlyLeuGlyAla 16

Db 64 ACCGAGCAGCGCATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

## RESULT 2

US-09-548-367D-50

Sequence 50, Application US/09548367D

Patent No. 6440698

GENERAL INFORMATION:

APPLICANT: GURNEY ET AL.

TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES

FILE REFERENCE: 29915/6280H

CURRENT FILING DATE: 2000-04-12

PRIOR APPLICATION NUMBER: US/09/548,367D

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 09/404,133

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: PCT/US99/20881

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 60/101,594

PRIOR FILING DATE: 1998-09-24

NUMBER OF SEQ ID NOS: 73

SOFTWARE: PatentIn version 3.1

SEQ ID NO 50

LENGTH: 1287

TYPE: DNA

ORGANISM: Artificial sequence

FEATURE:

OTHER INFORMATION: Hu-Asp2 (b) delta TM

US-09-548-367D-50

Alignment Scores:

Pred. No.:	1,48e-06	Length:	1287
Score:	16.00	Matches:	16
Percent Similarity:	100.00%	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	100.00%	Indels:	0
DB:	3	Gaps:	0

US-10-726-967A-3 (1-16) x US-09-548-367D-50 (1-1287)

QY 1 ThrglnhsglylIleArgLeuProLeuArgSerGlyLeuGlyAla 16

Db 64 ACCGAGCAGCGCATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

## RESULT 3

US-09-551-853D-50

Sequence 50, Application US/09551853D

Patent No. 6500667

GENERAL INFORMATION:

APPLICANT: GURNEY ET AL.

TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES

FILE REFERENCE: 29915/6280L

CURRENT FILING DATE: US/09/551,853D

PRIOR APPLICATION NUMBER: 2000-04-18

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 09/404,133

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: PCT/US99/20881

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 60/101,594

PRIOR FILING DATE: 1998-09-24

NUMBER OF SEQ ID NOS: 73

SOFTWARE: PatentIn version 3.1

SEQ ID NO 50

LENGTH: 1287

TYPE: DNA

ORGANISM: Artificial sequence

FEATURE:

OTHER INFORMATION: Hu-Asp2 (b) delta TM

US-09-551-853D-50

Alignment Scores:

Pred. No.:	1,48e-06	Length:	1287
Score:	16.00	Matches:	16
Percent Similarity:	100.00%	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	100.00%	Indels:	0
DB:	4	Gaps:	0

US-10-726-967A-3 (1-16) x US-09-551-853D-50 (1-1287)

QY 1 ThrglnhsglylIleArgLeuProLeuArgSerGlyLeuGlyAla 16

Db 64 ACCGAGCAGCGCATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

## RESULT 4

US-09-416-901B-50

Sequence 50, Application US/09416901B

Patent No. 6699671

GENERAL INFORMATION:

APPLICANT: GURNEY ET AL.

TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES

FILE REFERENCE: 29915/6280A

CURRENT FILING DATE: US/09/416,901B

PRIOR APPLICATION NUMBER: 1999-10-13

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 60/155,493

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 09/404,133

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: PCT/US99/20881

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 60/101,594

PRIOR FILING DATE: 1998-09-24

NUMBER OF SEQ ID NOS: 72

SOFTWARE: PatentIn version 3.1

SEQ ID NO 50

LENGTH: 1287

TYPE: DNA

ORGANISM: Artificial sequence

FEATURE:

OTHER INFORMATION: Hu-Asp2 (b) delta TM

US-09-416-901B-50

Alignment Scores:

Pred. No.:	1,48e-06	Length:	1287
Score:	16.00	Matches:	16
Percent Similarity:	100.00%	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	100.00%	Indels:	0
DB:	4	Gaps:	0

US-10-726-967A-3 (1-16) x US-09-416-901B-50 (1-1287)

QY 1 ThrglnhsglylIleArgLeuProLeuArgSerGlyLeuGlyAla 16

Db 64 ACCGAGCAGCGCATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

## RESULT 5

US-09-548-376D-50

Sequence 50, Application US/09548376D

Patent No. 6706485

GENERAL INFORMATION:

APPLICANT: GURNEY ET AL.

TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR

FILE REFERENCE: 29915/6280F

```

CURRENT APPLICATION NUMBER: US/09/548,376D
CURRENT FILING DATE: 2000-04-12
PRIOR APPLICATION NUMBER: US 60/155,493
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: US 09/404,133
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: PCT/US99/20881
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: US 60/101,594
PRIOR FILING DATE: 1998-09-24
NUMBER OF SEQ ID NOS: 73
SOFTWARE: PatentIn version 3.1
SEQ ID NO 50
LENGTH: 1287
TYPE: DNA
ORGANISM: Artificial sequence
FEATURE:
OTHER INFORMATION: Hu-Asp2(b) delta TM
US-09-548-376D-50

Alignment Scores:
Pred. No.: 1,48e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 4 Gaps: 0

US-10-726-967A-3 (1-16) x US-09-548-376D-50 (1-1287)
QY 1 ThrglnHieGlyIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16
Db 64 ACCGACGACGGCATCCGGCTGCCCTGCCGACGGCGCTGGGGGGCCGCC 111

RESULT 6
US-09-794-927A-50
Sequence 50, Application US/09794927A
Patent No. 6727074
GENERAL INFORMATION:
APPLICANT: Gurney et al.
TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses
FILE REFERENCE: 29915/6280FG
CURRENT APPLICATION NUMBER: US/09/794,927A
CURRENT FILING DATE: 2001-02-27
PRIOR APPLICATION NUMBER: 09/416,901
PRIOR FILING DATE: 1999-10-13
PRIOR APPLICATION NUMBER: 60/155,493
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: 09/404,133
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: PCT/US99/20881
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: 60/101,594
PRIOR FILING DATE: 1998-09-24
NUMBER OF SEQ ID NOS: 74
SOFTWARE: PatentIn Ver. 2.0
SEQ ID NO 50
LENGTH: 1287
TYPE: DNA
ORGANISM: Artificial sequence
FEATURE:
OTHER INFORMATION: Hu-Asp2(b) delta TM
US-09-794-927A-50

Alignment Scores:
Pred. No.: 1,48e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 4 Gaps: 0
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```

US-10-726-967A-3 (1-16) x US-09-794-927A-50 (1-1287)
QY 1 ThrglnHieGlyIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16
Db 64 ACCGACGACGGCATCCGGCTGCCCTGCCGACGGCGCTGGGGGGCCGCC 111

RESULT 7
US-09-548-373D-50
Sequence 50, Application US/09548373D
Patent No. 6737510
GENERAL INFORMATION:
APPLICANT: GURNEY ET AL.
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES
FILE REFERENCE: 29915/6280B
CURRENT APPLICATION NUMBER: US/09/548,373D
CURRENT FILING DATE: 2000-04-12
PRIOR APPLICATION NUMBER: US 60/155,493
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: US 09/404,133
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: PCT/US99/20881
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: US 60/101,594
PRIOR FILING DATE: 1998-09-24
NUMBER OF SEQ ID NOS: 73
SOFTWARE: PatentIn version 3.1
SEQ ID NO 50
LENGTH: 1287
TYPE: DNA
ORGANISM: Artificial sequence
FEATURE:
OTHER INFORMATION: Hu-Asp2(b) delta TM
US-09-548-373D-50

Alignment Scores:
Pred. No.: 1,48e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 4 Gaps: 0

US-10-726-967A-3 (1-16) x US-09-548-373D-50 (1-1287)
QY 1 ThrglnHieGlyIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16
Db 64 ACCGACGACGGCATCCGGCTGCCCTGCCGACGGCGCTGGGGGGCCGCC 111

RESULT 8
US-09-795-847B-50
Sequence 50, Application US/09795847B
Patent No. 6753163
GENERAL INFORMATION:
APPLICANT: Gurney, Mark E.
APPLICANT: Bienkowski, Michael J.
APPLICANT: Heinrikson, Robert L.
APPLICANT: Parodi, Luis A.
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
FILE REFERENCE: 28341/6280DE
CURRENT APPLICATION NUMBER: US/09/795,847B
CURRENT FILING DATE: 2001-02-28
PRIOR APPLICATION NUMBER: 09/416,901
PRIOR FILING DATE: 1999-10-13
PRIOR APPLICATION NUMBER: 60/155,493
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: 09/404,133
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: PCT/US99/20881
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: 60/101,594
PRIOR FILING DATE: 1999-09-23
PRIOR APPLICATION NUMBER: 60/101,594
```

PRIOR FILING DATE: 1998-09-24  
NUMBER OF SEQ ID NOS: 74  
SOFTWARE: Patentin Ver. 2.0  
SEQ ID NO 50  
LENGTH: 1287  
TYPE: DNA  
ORGANISM: Artificial sequence  
FEATURE:  
OTHER INFORMATION: Hu-Asp2 (b) delta TM  
US-09-795-847B-50

Alignment Scores:  
Pred. No.: 1,48e-06 Length: 1287  
Score: 16.00 Matches: 16  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 4 Gaps: 0

US-10-726-967A-3 (1-16) x US-09-795-847B-50 (1-1287)

Qy 1 ThrglnhsglyileargleuProleuArgserglyleuglyala 16  
Db 64 ACCGAGCAGGCAATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

RESULT 9  
US-09-869-414-50  
Sequence 50, Application US/09869414  
Patent No. 6790610  
GENERAL INFORMATION:  
APPLICANT: Belkowsky et al.  
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES  
FILE REFERENCE: 28341/6280M  
CURRENT APPLICATION NUMBER: US/09/869,414  
CURRENT FILING DATE: 2001-06-27  
PRIOR APPLICATION NUMBER: 09/416,901  
PRIOR FILING DATE: 1999-10-13  
PRIOR APPLICATION NUMBER: 60/155,493  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: 09/404,133  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: PCT/US99/20881  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: 60/101,594  
PRIOR FILING DATE: 1998-09-24  
NUMBER OF SEQ ID NOS: 73  
SOFTWARE: Patentin Ver. 2.0  
SEQ ID NO 50  
LENGTH: 1287  
TYPE: DNA  
ORGANISM: Artificial Sequence  
FEATURE:  
OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)  
US-09-869-414-50

Alignment Scores:  
Pred. No.: 1,48e-06 Length: 1287  
Score: 16.00 Matches: 16  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 4 Gaps: 0

US-10-726-967A-3 (1-16) x US-09-869-414-50 (1-1287)

Qy 1 ThrglnhsglyileargleuProleuArgserglyleuglyala 16  
Db 64 ACCGAGCAGGCAATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

RESULT 10  
US-09-548-366F-50

Sequence 50, Application US/09548366F  
Patent No. 6797487  
GENERAL INFORMATION:  
APPLICANT: GURNEY ET AL.  
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES  
FILE REFERENCE: 29915/6280J  
CURRENT APPLICATION NUMBER: US/09/548,366F  
CURRENT FILING DATE: 2000-04-12  
PRIOR APPLICATION NUMBER: US 60/155,493  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: US 09/404,133  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: PCT/US99/20881  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: US 60/101,594  
PRIOR FILING DATE: 1998-09-24  
NUMBER OF SEQ ID NOS: 73  
SOFTWARE: Patentin version 3.1  
SEQ ID NO 50  
LENGTH: 1287  
TYPE: DNA  
ORGANISM: Artificial sequence  
FEATURE:  
OTHER INFORMATION: Hu-Asp2 (b) delta TM  
US-09-548-366D-50

Alignment Scores:  
Pred. No.: 1,48e-06 Length: 1287  
Score: 16.00 Matches: 16  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 4 Gaps: 0

US-10-726-967A-3 (1-16) x US-09-548-366F-50 (1-1287)

Qy 1 ThrglnhsglyileargleuProleuArgserglyleuglyala 16  
Db 64 ACCGAGCAGGCAATCCGGCTGCCCTGCGCAGCGGCTGGGGGGCGCC 111

RESULT 11  
US-09-548-368D-50  
Sequence 50, Application US/09548368D  
Patent No. 6825023  
GENERAL INFORMATION:  
APPLICANT: GURNEY ET AL.  
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES  
FILE REFERENCE: 29915/6280C  
CURRENT APPLICATION NUMBER: US/09/548,368D  
CURRENT FILING DATE: 2000-04-12  
PRIOR APPLICATION NUMBER: US 60/155,493  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: US 09/404,133  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: PCT/US99/20881  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: US 60/101,594  
PRIOR FILING DATE: 1998-09-24  
NUMBER OF SEQ ID NOS: 73  
SOFTWARE: Patentin version 3.1  
SEQ ID NO 50  
LENGTH: 1287  
TYPE: DNA  
ORGANISM: Artificial sequence  
FEATURE:  
OTHER INFORMATION: Hu-Asp2 (b) delta TM  
US-09-548-368D-50

Alignment Scores:  
Pred. No.: 1,48e-06 Length: 1287  
Score: 16.00 Matches: 16



Percent Similarity: 100.00%      Conservative: 0  
Best Local Similarity: 100.00%      Mismatches: 0  
Query Match: 100.00%      Indels: 0  
DB: 4      Gaps: 0

US-10-726-967A-3 (1-16) x US-09-548-368D-50 (1-1287)

QY 1 ThGlnHIGLYIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16  
|||||  
DB 64 ACCCAGCAGCGCATCCGGCTGCCCTCGCCAGCGGCTGGGGGGCCGCC 111

RESULT 12  
US-09-794-925A-50

Sequence 50, Application US/09794925A

Patent No. 6828117

GENERAL INFORMATION:

APPLICANT: Gurney et al.

TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses

FILE REFERENCE: 29915/6280H1

CURRENT APPLICATION NUMBER: US/09/794,925A

CURRENT FILING DATE: 2001-02-27

PRIOR APPLICATION NUMBER: 09/416,901

PRIOR FILING DATE: 1999-10-13

PRIOR APPLICATION NUMBER: 60/155,493

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: 09/404,133

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: PCT/US99/20881

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: 60/101,594

PRIOR FILING DATE: 1998-09-24

NUMBER OF SEQ ID NOS: 74

SOFTWARE: PatentIn Ver. 2.0

SEQ ID NO 50

LENGTH: 1287

TYPE: DNA

ORGANISM: Artificial sequence

FEATURE: OTHER INFORMATION: Hu-Amp2 (b) delta TM

US-09-794-925A-50

Alignment Scores:

Pred. No.: 1,48e-06      Length: 1287

Score: 16.00      Matches: 16

Percent Similarity: 100.00%      Conservative: 0

Best Local Similarity: 100.00%      Mismatches: 0

Query Match: 100.00%      Indels: 0

DB: 4      Gaps: 0

US-10-726-967A-3 (1-16) x US-09-794-925A-50 (1-1287)

QY 1 ThGlnHIGLYIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16  
|||||  
DB 64 ACCCAGCAGCGCATCCGGCTGCCCTCGCCAGCGGCTGGGGGGCCGCC 111

RESULT 13

US-09-548-372D-25

Sequence 25, Application US/09548372D

Patent No. 6420534

GENERAL INFORMATION:

APPLICANT: Gurney et al.

TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES

FILE REFERENCE: 29915/62801

CURRENT APPLICATION NUMBER: US/09/548,372D

CURRENT FILING DATE: 2000-04-12

PRIOR APPLICATION NUMBER: US 60/155,493

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 09/404,133

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: PCT/US99/20881

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 60/101,594  
PRIOR FILING DATE: 1998-09-24  
NUMBER OF SEQ ID NOS: 73  
SOFTWARE: PatentIn version 3.1  
SEQ ID NO 25  
LENGTH: 1302  
TYPE: DNA  
ORGANISM: Homo sapiens  
US-09-548-372D-25

Alignment Scores:

Pred. No.: 1.5e-06      Length: 1302

Score: 16.00      Matches: 16

Percent Similarity: 100.00%      Conservative: 0

Best Local Similarity: 100.00%      Mismatches: 0

Query Match: 100.00%      Indels: 0

DB: 3      Gaps: 0

US-10-726-967A-3 (1-16) x US-09-548-372D-25 (1-1302)

QY 1 ThGlnHIGLYIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16  
|||||  
DB 4 ACTCAGCATGTATTGCTCTGCCACTGCTAGCGGCTGGGTGGTCT 51

RESULT 14

US-09-548-367D-25

Sequence 25, Application US/09548367D

Patent No. 6440698

GENERAL INFORMATION:

APPLICANT: Gurney et al.

TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES

FILE REFERENCE: 29915/6280H

CURRENT APPLICATION NUMBER: US/09/548,367D

CURRENT FILING DATE: 2000-04-12

PRIOR APPLICATION NUMBER: US 60/155,493

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 09/404,133

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: PCT/US99/20881

PRIOR FILING DATE: 1999-09-23

PRIOR APPLICATION NUMBER: US 60/101,594

PRIOR FILING DATE: 1998-09-24

NUMBER OF SEQ ID NOS: 73

SOFTWARE: PatentIn version 3.1

SEQ ID NO 25

LENGTH: 1302

TYPE: DNA

ORGANISM: Homo sapiens

US-09-548-367D-25

Alignment Scores:

Pred. No.: 1.5e-06      Length: 1302

Score: 16.00      Matches: 16

Percent Similarity: 100.00%      Conservative: 0

Best Local Similarity: 100.00%      Mismatches: 0

Query Match: 100.00%      Indels: 0

DB: 3      Gaps: 0

US-10-726-967A-3 (1-16) x US-09-548-367D-25 (1-1302)

QY 1 ThGlnHIGLYIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16  
|||||  
DB 4 ACTCAGCATGTATTGCTCTGCCACTGCTAGCGGCTGGGTGGTCT 51

RESULT 15

US-09-551-853D-25

Sequence 25, Application US/09551853D

Patent No. 650667

GENERAL INFORMATION:

APPLICANT: Gurney et al.

TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES

FILE REFERENCE: 29915/62801

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1 FILE REFERENCE: 29915/6280L
2 CURRENT APPLICATION NUMBER: US/09/551,853D
3 PRIOR FILING DATE: 2000-04-18
4 PRIOR APPLICATION NUMBER: US 60/155,493
5 PRIOR FILING DATE: 1999-09-23
6 PRIOR APPLICATION NUMBER: US 09/404,133
7 PRIOR FILING DATE: 1999-09-23
8 PRIOR APPLICATION NUMBER: PCT/US99/20881
9 PRIOR FILING DATE: 1999-09-23
10 PRIOR APPLICATION NUMBER: US 60/101,594
11 PRIOR FILING DATE: 1998-09-24
12 NUMBER OF SEQ ID NOS: 73
13 SOFTWARE: PatencIn version 3.1
14 SEQ ID NO 25
15 LENGTH: 1302
16 TYPE: DNA
17 ORGANISM: Homo sapiens
18
19 US-09-551-853D-25

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## OM protein - nucleic search, using frame\_plus\_p2n model

Run on: July 27, 2005, 19:03:29 ; Search time 625 Seconds

(without alignments)  
165.537 Million cell updates/sec

Title: US-10-726-967a-3

Perfect score: 16

Sequence: 1 TQHGRLPLRSLGCA 16

## Scoring table:

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	Ygapop 60.0	Ygapext 60.0
	Fgapop 6.0	Fgapext 7.0
	Delop 6.0	Delext 7.0

Searched: 7277826 seqs, 323139505 residues

Word size: 1

Total number of hits satisfying chosen parameters: 14539411

Minimum DB seq length: 0

Maximum DB seq length: 200000000

Post-processing: listing first 45 summaries

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-FAPOP=6 -FAPEXT=7 -YGAPOP=60 -YGAPEXT=60 -DBLOP=6 -DELEXT=7

## Database :

Published Applications NA:\*

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- 2: /cgn2\_6/ptodata/1/pubpna/PCT\_NEW\_PUB.seq:\*
- 3: /cgn2\_6/ptodata/1/pubpna/US06\_NEW\_PUB.seq:\*
- 4: /cgn2\_6/ptodata/1/pubpna/US06\_PUBCOMB.seq:\*
- 5: /cgn2\_6/ptodata/1/pubpna/US07\_NEW\_PUB.seq:\*
- 6: /cgn2\_6/ptodata/1/pubpna/PCTUS\_PUBCOMB.seq:\*
- 7: /cgn2\_6/ptodata/1/pubpna/US08\_NEW\_PUB.seq:\*
- 8: /cgn2\_6/ptodata/1/pubpna/US08\_PUBCOMB.seq:\*
- 9: /cgn2\_6/ptodata/1/pubpna/US09A\_PUBCOMB.seq:\*
- 10: /cgn2\_6/ptodata/1/pubpna/US09B\_PUBCOMB.seq:\*
- 11: /cgn2\_6/ptodata/1/pubpna/US09C\_PUBCOMB.seq:\*
- 12: /cgn2\_6/ptodata/1/pubpna/US09\_NEW\_PUB.seq:\*
- 13: /cgn2\_6/ptodata/1/pubpna/US10A\_PUBCOMB.seq:\*
- 14: /cgn2\_6/ptodata/1/pubpna/US10B\_PUBCOMB.seq:\*
- 15: /cgn2\_6/ptodata/1/pubpna/US10D\_PUBCOMB.seq:\*
- 16: /cgn2\_6/ptodata/1/pubpna/US10E\_PUBCOMB.seq:\*
- 17: /cgn2\_6/ptodata/1/pubpna/US10F\_PUBCOMB.seq:\*
- 18: /cgn2\_6/ptodata/1/pubpna/US10G\_PUBCOMB.seq:\*
- 19: /cgn2\_6/ptodata/1/pubpna/US10H\_PUBCOMB.seq:\*
- 20: /cgn2\_6/ptodata/1/pubpna/US10I\_PUBCOMB.seq:\*
- 21: /cgn2\_6/ptodata/1/pubpna/US10J\_PUBCOMB.seq:\*
- 22: /cgn2\_6/ptodata/1/pubpna/US11\_NEW\_PUB.seq:\*
- 23: /cgn2\_6/ptodata/1/pubpna/US11\_PUBCOMB.seq:\*
- 24: /cgn2\_6/ptodata/1/pubpna/US11\_NEW\_PUB.seq:\*
- 25: /cgn2\_6/ptodata/1/pubpna/US60\_NEW\_PUB.seq:\*
- 26: /cgn2\_6/ptodata/1/pubpna/US60\_PUBCOMB.seq:\*

Pred. No. is the number of results predicted by chance to have a

score greater than or equal to the score of the result being printed,  
and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	DB	ID	Description
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2	16	100.0	1287	9	US-09-795-847-50	Sequence 50, Appl
3	16	100.0	1287	9	US-09-794-743-50	Sequence 50, Appl
4	16	100.0	1287	9	US-09-794-748-50	Sequence 50, Appl
5	16	100.0	1287	9	US-09-794-925-50	Sequence 50, Appl
6	16	100.0	1287	9	US-09-681-442-50	Sequence 50, Appl
7	16	100.0	1287	10	US-09-869-414-50	Sequence 50, Appl
8	16	100.0	1287	10	US-09-548-366-50	Sequence 50, Appl
9	16	100.0	1287	18	US-10-652-927-50	Sequence 50, Appl
10	16	100.0	1287	18	US-10-652-830-50	Sequence 50, Appl
11	16	100.0	1287	19	US-10-652-045-50	Sequence 50, Appl
12	16	100.0	1287	20	US-10-476-935-50	Sequence 50, Appl
13	16	100.0	1287	21	US-10-477-076-50	Sequence 50, Appl
14	16	100.0	1302	9	US-09-794-927-25	Sequence 25, Appl
15	16	100.0	1302	9	US-09-795-847-25	Sequence 25, Appl
16	16	100.0	1302	9	US-09-794-743-25	Sequence 25, Appl
17	16	100.0	1302	9	US-09-794-748-25	Sequence 25, Appl
18	16	100.0	1302	9	US-09-794-925-25	Sequence 25, Appl
19	16	100.0	1302	9	US-09-681-442-25	Sequence 25, Appl
20	16	100.0	1302	10	US-09-869-414-25	Sequence 25, Appl
21	16	100.0	1302	10	US-09-548-366-25	Sequence 25, Appl
22	16	100.0	1302	18	US-10-652-927-25	Sequence 25, Appl
23	16	100.0	1302	18	US-10-652-830-25	Sequence 25, Appl
24	16	100.0	1302	19	US-10-652-045-25	Sequence 25, Appl
25	16	100.0	1302	20	US-10-476-935-25	Sequence 25, Appl
26	16	100.0	1302	20	US-10-477-076-25	Sequence 25, Appl
27	16	100.0	1302	21	US-10-477-076-25	Sequence 25, Appl
28	16	100.0	1305	9	US-09-794-927-52	Sequence 52, Appl
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30	16	100.0	1305	9	US-09-794-743-52	Sequence 52, Appl
31	16	100.0	1305	9	US-09-794-748-52	Sequence 52, Appl
32	16	100.0	1305	9	US-09-794-925-52	Sequence 52, Appl
33	16	100.0	1305	9	US-09-681-442-52	Sequence 52, Appl
34	16	100.0	1305	10	US-09-869-414-52	Sequence 52, Appl
35	16	100.0	1305	10	US-09-548-366-52	Sequence 52, Appl
36	16	100.0	1305	18	US-10-652-927-52	Sequence 52, Appl
37	16	100.0	1305	18	US-10-652-830-52	Sequence 52, Appl
38	16	100.0	1305	19	US-10-652-045-52	Sequence 52, Appl
39	16	100.0	1305	20	US-10-476-935-52	Sequence 52, Appl
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42	16	100.0	1341	9	US-09-795-847-21	Sequence 21, Appl
43	16	100.0	1341	9	US-09-794-743-21	Sequence 21, Appl
44	16	100.0	1341	9	US-09-794-748-21	Sequence 21, Appl
45	16	100.0	1341	9	US-09-794-925-21	Sequence 21, Appl

## ALIGNMENTS

RESULT 1  
US-09-794-927-50  
; Sequence 50, Application US/09794927  
; Patent No. US20010016324A1  
GENERAL INFORMATION:  
; APPLICANT: Guiney, Mark E.  
; APPLICANT: Blenkowaki, Michael J.  
; APPLICANT: Heinrichson, Robert L.  
; APPLICANT: Parodi, Luis A.  
; APPLICANT: Yan, Riqiang  
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND  
; TITLE OF INVENTION: THEREFOR  
; FILE REFERENCE: 28341/6280FG  
; CURRENT APPLICATION NUMBER: US/09/794, 927  
; CURRENT FILING DATE: 2001-02-27

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; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
; US-09-794-927-50

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Alignment Scores:
Pred. No.: 2,94e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0

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US-10-726-967a-3 (1-16) x US-09-794-927-50 (1-1287)

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QY 1 ThrglnHsglyYlleArgLeuProLeuAArgSerGlyLeuGlyGlyAla 16
Db 64 ACCCAGCAGCGCATCCGCTGCCCTCGCCAGCGCGCTCGGGGGGCC 111

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#### RESULT 2

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; Sequence 50, Application US/09795847
; Patent No. US20010018208A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrichson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND
; TITLE OF INVENTION: USES
; FILE REFERENCE: 28341/6280DE
; CURRENT APPLICATION NUMBER: US/09/795,847
; CURRENT FILING DATE: 2001-02-28
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
; US-09-795-847-50

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#### Alignment Scores:

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Pred. No.: 2,94e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0

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US-10-726-967a-3 (1-16) x US-09-795-847-50 (1-1287)

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QY 1 ThrglnHsglyYlleArgLeuProLeuAArgSerGlyLeuGlyGlyAla 16
Db 64 ACCCAGCAGCGCATCCGCTGCCCTCGCCAGCGCGCTCGGGGGGCC 111

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#### RESULT 3

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; Sequence 50, Application US/09794743
; Patent No. US20010021391A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrichson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND
; TITLE OF INVENTION: USES
; FILE REFERENCE: 28341/6280BC
; CURRENT APPLICATION NUMBER: US/09/794,743
; CURRENT FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
; US-09-794-743-50

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Alignment Scores:
Pred. No.: 2,94e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0

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US-10-726-967a-3 (1-16) x US-09-794-743-50 (1-1287)

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QY 1 ThrglnHsglyYlleArgLeuProLeuAArgSerGlyLeuGlyGlyAla 16
Db 64 ACCCAGCAGCGCATCCGCTGCCCTCGCCAGCGCGCTCGGGGGGCC 111

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#### RESULT 4

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; Sequence 50, Application US/09794748
; Patent No. US20020037315A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrichson, Robert L.
; APPLICANT: Parodi, Luis A.

```

```

; APPLICANT: Yan, Rigdang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND
; TITLE OF INVENTION: USES
; TITLE OF INVENTION: THEREFOR
; FILE REFERENCE: 28341/6280JL
; CURRENT APPLICATION NUMBER: US/09/794,748
; CURRENT FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Ap2 (b)
; US-09-794-748-50

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Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
Gaps: 0
DB: 9

US-10-726-967a-3 (1-16) x US-09-794-748-50 (1-1287)

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Db 64 ACCGACGACGCGATCCGCGCTGCGCCCTGCGCAGCGCGCTGGGGGCGGCC 111

RESULT 5
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; Sequence 50, Application US/09794925
; Patent No. US20020064819A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowiak, Michael J.
; APPLICANT: Heinrikson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Rigdang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; TITLE OF INVENTION: THEREFOR
; FILE REFERENCE: 28341/6280HI
; CURRENT APPLICATION NUMBER: US/09/794,925
; CURRENT FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
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; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Ap2 (b)
; OTHER INFORMATION: delta TM
; US-09-794-925-50

Alignment Scores:
Pred. No.: 2,94e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
Gaps: 0
DB: 9

US-10-726-967a-3 (1-16) x US-09-794-925-50 (1-1287)

Qy 1 ThrglnHieGlyYleArGleuPProleuAArgSerGlyLeuGlyYAla 16
Db 64 ACCGACGACGCGATCCGCGCTGCGCCCTGCGCAGCGCGCTGGGGGCGGCC 111

RESULT 6
US-09-681-442-50
; Sequence 50, Application US/09681442
; Patent No. US20020081634A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowiak, Michael J.
; APPLICANT: Heinrikson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Rigdang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; TITLE OF INVENTION: THEREFOR
; FILE REFERENCE: 28341/6280FG
; CURRENT APPLICATION NUMBER: US/09/681,442
; CURRENT FILING DATE: 2001-04-05
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Ap2 (b)
; OTHER INFORMATION: delta TM
; US-09-681-442-50

Alignment Scores:
Pred. No.: 2,94e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
Gaps: 0
DB: 9

US-10-726-967a-3 (1-16) x US-09-681-442-50 (1-1287)

Qy 1 ThrglnHieGlyYleArGleuPProleuAArgSerGlyLeuGlyYAla 16
Db 64 ACCGACGACGCGATCCGCGCTGCGCCCTGCGCAGCGCGCTGGGGGCGGCC 111

RESULT 7
US-09-869-414-50
; Sequence 50, Application US/09869414
; Publication No. US20030077226A1
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; GENERAL INFORMATION:
; APPLICANT: Bienkowski et al.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; FILE REFERENCE: 28341/6280M
; CURRENT APPLICATION NUMBER: US/09/869,414
; PRIOR FILING DATE: 2001-06-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentln Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2(b)
US-09-869-414-50

Alignment Scores:
Pred. No.: 2,94e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 10 Gaps: 0

US-10-726-967A-3 (1-16) x US-09-869-414-50 (1-1287)
QY 1 Thrglnhlglytlleargleuproleuargserglyleuglyala 16
Db 64 ACCGAGCAGCGCATCCGGCTGCCCTCGCGAGCGGCTGGGGGGCGCC 111

RESULT 8
US-09-548-366-50
; Sequence 50, Application US/09548366
; Publication No. US20030104365A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrichson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND
; FILE REFERENCE: 28341/6280A
; CURRENT APPLICATION NUMBER: US/09/548,366
; PRIOR FILING DATE: 2000-04-12
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 65
; SOFTWARE: Patentln Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2(b)
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; OTHER INFORMATION: delta TM
US-09-548-366-50

Alignment Scores:
Pred. No.: 2,94e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 10 Gaps: 0

US-10-726-967A-3 (1-16) x US-09-548-366-50 (1-1287)
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Db 64 ACCGAGCAGCGCATCCGGCTGCCCTCGCGAGCGGCTGGGGGGCGCC 111

RESULT 9
US-10-652-927-50
; Sequence 50, Application US/10652927
; Publication No. US20040043408A1
; GENERAL INFORMATION:
; APPLICANT: Gurney et al.
; TITLE OF INVENTION: Alzheimer's Disease Secretase, App Substrates Therefor and Uses
; FILE REFERENCE: 28915/6280N3
; CURRENT APPLICATION NUMBER: US/10/652,927
; PRIOR FILING DATE: 2003-08-29
; PRIOR APPLICATION NUMBER: 09/794,925
; PRIOR FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 74
; SOFTWARE: Patentln Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial sequence
; FEATURE:
; OTHER INFORMATION: Hu-Asp2(b) delta TM
US-10-652-927-50

Alignment Scores:
Pred. No.: 2,94e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 18 Gaps: 0

US-10-726-967A-3 (1-16) x US-10-652-927-50 (1-1287)
QY 1 Thrglnhlglytlleargleuproleuargserglyleuglyala 16
Db 64 ACCGAGCAGCGCATCCGGCTGCCCTCGCGAGCGGCTGGGGGGCGCC 111

RESULT 10
US-10-652-830-50
; Sequence 50, Application US/10652830
; Publication No. US20040048303A1
; GENERAL INFORMATION:
; APPLICANT: Gurney et al.
; TITLE OF INVENTION: Alzheimer's Disease Secretase, App Substrates Therefor and Uses
; FILE REFERENCE: 29915/6280N1
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; CURRENT APPLICATION NUMBER: US/10/652,830
; CURRENT FILING DATE: 2003-08-29
; PRIOR APPLICATION NUMBER: 09/794,925
; PRIOR FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 74
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial sequence
; FEATURE:
; OTHER INFORMATION: Hu-Asp2 (b) delta TM
US-10-652-830-50

Alignment Scores:
Pred. No.:      2,94e-06      Length:      1287
Score:          16.00         Matches:      16
Percent Similarity: 100.00%    Conservative: 0
Best Local Similarity: 100.00% Mismatches:      0
Query Match:    100.00%       Indels:        0
DB:             18           Gaps:          0

US-10-726-967a-3 (1-16) x US-10-652-830-50 (1-1287)

QY      1  ThrGlnHieGlyIleArgLeuProLeuArgSerGlyLeuGlyAla 16
Db      64 ACCGACGACGGCATCCGCTGCCCTCGCCACGCGGCTCGGGGGGCC 111

RESULT 11
US-10-652-045-50
; Sequence 50, Application US/10652045
; Publication No. US20040166507A1
; GENERAL INFORMATION:
; APPLICANT: Gurney et al.
; TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses
; FILE REFERENCE: 29915/6280M2
; CURRENT APPLICATION NUMBER: US/10/652,045
; CURRENT FILING DATE: 2003-08-29
; PRIOR APPLICATION NUMBER: 09/794,925
; PRIOR FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 74
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial sequence
; FEATURE:
; OTHER INFORMATION: Hu-Asp2 (b) delta TM
US-10-652-045-50

Alignment Scores:
Pred. No.:      2,94e-06      Length:      1287
Score:          16.00         Matches:      16
Percent Similarity: 100.00%    Conservative: 0
Best Local Similarity: 100.00% Mismatches:      0
Query Match:    100.00%       Indels:        0
DB:             18           Gaps:          0

US-10-726-967a-3 (1-16) x US-10-652-830-50 (1-1287)

QY      1  ThrGlnHieGlyIleArgLeuProLeuArgSerGlyLeuGlyAla 16
Db      64 ACCGACGACGGCATCCGCTGCCCTCGCCACGCGGCTCGGGGGGCC 111
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Score:          16.00         Matches:      16
Percent Similarity: 100.00%    Conservative: 0
Best Local Similarity: 100.00% Mismatches:      0
Query Match:    100.00%       Indels:        0
DB:             19           Gaps:          0

US-10-726-967a-3 (1-16) x US-10-652-045-50 (1-1287)

QY      1  ThrGlnHieGlyIleArgLeuProLeuArgSerGlyLeuGlyAla 16
Db      64 ACCGACGACGGCATCCGCTGCCCTCGCCACGCGGCTCGGGGGGCC 111

RESULT 12
US-10-476-935-50
; Sequence 50, Application US/10476935
; Publication No. US20040234976A1
; GENERAL INFORMATION:
; APPLICANT: Beinkowski et al.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; FILE REFERENCE: 28341/6280M1
; CURRENT APPLICATION NUMBER: US/10/476,935
; CURRENT FILING DATE: 2003-11-06
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
US-10-476-935-50

Alignment Scores:
Pred. No.:      2,94e-06      Length:      1287
Score:          16.00         Matches:      16
Percent Similarity: 100.00%    Conservative: 0
Best Local Similarity: 100.00% Mismatches:      0
Query Match:    100.00%       Indels:        0
DB:             20           Gaps:          0

US-10-726-967a-3 (1-16) x US-10-476-935-50 (1-1287)

QY      1  ThrGlnHieGlyIleArgLeuProLeuArgSerGlyLeuGlyAla 16
Db      64 ACCGACGACGGCATCCGCTGCCCTCGCCACGCGGCTCGGGGGGCC 111

RESULT 13
US-10-477-076-50
; Sequence 50, Application US/10477076
; Publication No. US20050080232A1
; GENERAL INFORMATION:
; APPLICANT: Beinkowski et al.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; FILE REFERENCE: 28341/6280M2
; CURRENT APPLICATION NUMBER: US/10/477,076
; CURRENT FILING DATE: 2003-11-06
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
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; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentln Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
; OTHER INFORMATION: delta TM
US-10-477-076-50

Alignment Scores:
Pred. No.: 2.94e-06 Length: 1287
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 21 Gaps: 0

US-10-726-967a-3 (1-16) x US-10-477-076-50 (1-1287)

Qy 1 ThrglnhsglylleargleuProleuArgserglyleuglyYAla 16
Db 64 ACCGACGACGGCATCGGCTGCCCTGCGACGCGCGCTGGGGGGCGCC 111

RESULT 14
US-09-794-927-25
; Sequence 25, Application US/09794927
; Patent No. US20010016324A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heintikson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND
; TITLE OF INVENTION: USES
; FILE REFERENCE: 28341/6280FG
; CURRENT APPLICATION NUMBER: US/09/794,927
; PRIOR FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentln Ver. 2.0
; SEQ ID NO 25
; LENGTH: 1302
; TYPE: DNA
; ORGANISM: Homo sapiens
US-09-794-927-25

Alignment Scores:
Pred. No.: 2.97e-06 Length: 1302
Score: 16.00 Matches: 16
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0

US-10-726-967a-3 (1-16) x US-10-477-076-50 (1-1287)

Qy 1 ThrglnhsglylleargleuProleuArgserglyleuglyYAla 16
Db 64 ACCGACGACGGCATCGGCTGCCCTGCGACGCGCGCTGGGGGGCGCC 111

US-10-726-967a-3 (1-16) x US-09-795-847-25 (1-1302)

Qy 1 ThrglnhsglylleargleuProleuArgserglyleuglyYAla 16
Db 4 ACTGAGCATGGTATTCGTCCACTGCGTAGCGGCTCGGGTGCT 51

Search completed: July 27, 2005, 20:48:51
Job time : 626 secs
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GenCore version 5.1.6  
Copyright (c) 1993 - 2005 CompuGen Ltd.

OM protein - nucleic search, using frame\_plus\_p2n model

Run on: July 27, 2005, 18:50:24 ; Search time 3143 Seconds  
(without alignments)  
193.773 Million cell updates/sec

Title: US-10-726-967A-3

Perfect score: 16  
Sequence: 1 TGHGRLPLRSLGCA 16

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Ygapop 60.0 , Ygapext 60.0  
Fgapop 6.0 , Fgapext 7.0  
Delop 6.0 , Delext 7.0

Searched: 34239544 seqs, 19032134700 residues

Word size: 1

Total number of hits satisfying chosen parameters: 68473592

Minimum DB seq length: 0  
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Post-processing: listing first 45 summaries

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-O/cn2.1/USPTO.spool.p/US10726967/runat.26072005.130816.6487/app.query.fasta.1.199  
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-DOCALLIGN=200 -THR.SCORE=quality -THR.MIN=1 -ALIGN=15 -MODE=LOCAL -OUTFMT=prco  
-NORM=ext -HEAPSIZE=500 -MINLEN=0 -MAXLEN=2000000000  
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Database :

EST:  
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2: gb\_est2:  
3: gb\_hic:  
4: gb\_est3:  
5: gb\_est4:  
6: gb\_est5:  
7: gb\_est6:  
8: gb\_gse1:  
9: gb\_gse2:

Pred. No. is the number of results predicted by chance to have a  
score greater than or equal to the score of the result being printed,  
and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	16	100.0	611	7	CN484125 hw42d08.y
2	16	100.0	1506	9	AY417360 Homo sapi
3	13	81.2	424	4	B1337739
4	13	81.2	563	4	BG833894
5	13	81.2	616	5	BP452814
6	11	68.8	171	9	CE200035
7	10	62.5	493	7	CN692524
8	10	62.5	533	7	CN697484
9	10	62.5	547	7	CF903755

10	10	62.5	563	7	CF906581
11	10	62.5	576	7	CF171218
12	10	62.5	600	6	BY713879
13	10	62.5	619	2	BB644736
14	10	62.5	637	6	CD348605
15	10	62.5	639	2	BB632244
16	10	62.5	639	2	BB652612
17	10	62.5	640	6	BY724111
18	10	62.5	650	2	BB624169
19	10	62.5	653	6	BY727440
20	10	62.5	673	2	BB640442
21	10	62.5	673	2	BB650860
22	10	62.5	727	6	CA749486
23	10	62.5	836	5	BK374914
24	10	62.5	844	1	AL544727
25	10	62.5	1123	5	BX376891
26	10	62.5	1506	9	AY117362
27	10	62.5	3634	3	AK041285
28	10	62.5	3859	3	AK014464
29	10	62.5	3877	3	AK033112
30	10	62.5	4046	3	AK049626
31	10	62.5	4048	3	AK082317
32	10	62.5	4101	3	AK046175
33	10	56.2	409	6	CB215573
34	9	56.2	566	8	BH881298
35	9	56.2	913	9	CG310449
36	9	56.2	965	5	BX393934
37	9	56.2	1765	2	BF572300
38	9	56.2	156	7	CO060378
39	8	50.0	220	7	CF681698
40	8	50.0	234	2	BB569323
41	8	50.0	258	2	BF872950
42	8	50.0	272	4	BM258402
43	8	50.0	295	4	BM574145
44	8	50.0	305	9	CB810341
45	8	50.0			

#### ALIGNMENTS

RESULT 1  
CN484125 611 bp mRNA linear EST 26-APR-2004  
LOCUS hw42d08.y2 Human primary human ocular pericytes. Unamplified (hw)  
DEFINITION Homo sapiens CDNA clone hw42d08 5', mRNA sequence.  
ACCESSION CN484125  
VERSION CN484125.1 GI:46565629  
KEYWORDS EST.  
SOURCE Homo sapiens (human)  
ORGANISM Homo sapiens

REFERENCE Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Primates; Catarrhini; Homiidae; Homo.  
AUTHORS Tsai, J.Y. and Wistow, G.  
TITLE 1 (bases 1 to 611)  
PERCYTES

JOURNAL Expressed sequence tag analysis of cultured primary human ocular  
pericytes  
COMMENT Unpublished (2004)  
CONTACT: Wistow G  
SECTION ON MOLECULAR STRUCTURE AND FUNCTION  
NATIONAL EYE INSTITUTE  
6/331, NIH, Bethesda, MD 20892-2740, USA  
TEL: 301 402 3452  
FAX: 301 496 0078  
EMAIL: gwaeme@nei.nih.gov  
PLATE: 42 row: d column: 08  
SEG PRIMER: M13Rpl reverse primer (AB1).  
LOCATION/Qualifiers

#### FEATURES

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/organism="Homo sapiens"  
/mol\_type="mRNA"  
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/clone="hw42d08"  
/cell\_type="pericytes"

/dev\_stage="Adult"  
/lab\_host="EMDH10B"  
/clone\_lib="human primary human ocular pericytes.  
Unamplified (hw)"  
/note="Organ: Eye; Vector: pSPORT1; RNA was extracted from  
primary human pericytes in culture. A directionally cloned  
cDNA library in the pSPORT vector (Invitrogen) was  
constructed at Bioserve Biotechnology (Laurel MD)  
essentially following the protocols of the Superscript  
Plasmid System full details of which are contained in the  
manufacturer's instruction manual  
(http://www.lifetechn.com/). First strand synthesis was  
carried out using a Not I primer-adaptor  
[5'-pGACTAGTCTAGATCGAGCGCGCC(T)15-3']. cDNA was  
cloned in Not I/Sal I sites. EST analysis was performed at  
the NIH Intramural Sequencing Center (NISC)."

## ORIGIN

## Alignment Scores:

Pred. No.: 9,47e-06 Length: 611  
Score: 16.00 Matches: 16  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: Gaps: 0

US-10-726-967A-3 (1-16) x CM484125 (1-611)

QY 1 ThrGlnHtGlyIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16

Db 381 ACCGACGCGCATCCGCTGCCCTGCCAGCGGCTGGGGGCGCC 428

## RESULT 2

AY417360

LOCUS AY417360 1506 bp DNA linear GSS 17-DEC-2003  
DEFINITION Homo sapiens BACE gene, VIRUAL TRANSCRIPT, partial sequence.  
ACCESSION AY417360  
VERSION AY417360.1 GI:39773320  
KEYWORDS GSS.  
SOURCE Homo sapiens (human)  
ORGANISM Homo sapiens  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.  
1 (bases 1 to 1506)  
Clark,A.G., Gnanowski,S., Nielson,R., Thomas,P., Kejarival,A.,  
Todd,M.A., Tanenbaum,D.M., Civello,D.R., Lu,F., Murphy,B.,  
Ferreira,S., Wang,G., Zheng,X.H., White,T.J., Sninsky,J.J.,  
Adams,M.D. and Cargill,M.  
Inferring nonneutral evolution from human-chimp-mouse orthologous  
gene trios  
Science 302 (5652), 1960-1963 (2003)

JOURNAL PUBMED 14671302  
AUTHORS 2 (bases 1 to 1506)  
Clark,A.G., Gnanowski,S., Nielson,R., Thomas,P., Kejarival,A.,  
Todd,M.A., Tanenbaum,D.M., Civello,D.R., Lu,F., Murphy,B.,  
Ferreira,S., Wang,G., Zheng,X.H., White,T.J., Sninsky,J.J.,  
Adams,M.D. and Cargill,M.  
Direct Submision  
Submitted (16-NOV-2003) Celera Genomics, 45 West Gude Drive,  
Rockville, MD 20850, USA  
This sequence was made by sequencing genomic exons and ordering  
them based on alignment.  
Location/Qualifiers  
1..1506  
/organism="Homo sapiens"  
/mol\_type="genomic DNA"  
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/locus\_tag="HCM6198"

## ORIGIN

Alignment Scores:  
Pred. No.: 2,11e-05 Length: 1506  
Score: 16.00 Matches: 16  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: Gaps: 0

US-10-726-967A-3 (1-16) x AY417360 (1-1506)

QY 1 ThrGlnHtGlyIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16

Db 64 ACCGACGCGCATCCGCTGCCCTGCCAGCGGCTGGGGGCGCC 111

## RESULT 3

B1337739

LOCUS B1337739 424 bp mRNA linear EST 30-JUL-2001  
DEFINITION 361259 MARC 1P1G Sus scrofa cDNA 5', mRNA sequence.  
ACCESSION B1337739  
VERSION B1337739.1 GI:15031022  
KEYWORDS EST.  
SOURCE Sus scrofa (pig)  
ORGANISM Sus scrofa  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Cetartiodactyla; Suina; Suidae; Sus.  
1 (bases 1 to 424)  
Fahrenkrug,S.C., Smith,T.P.L., Freking,B.A., Cho,J., White,J.,  
Vallet,J., Wise,T., Rohrer,G.A., Pertea,G., Sultana,R.,  
Quackenbush,J. and Keefe,J.W.  
Porcine gene discovery by normalized cDNA-library sequencing and  
EST cluster assembly  
Mamm. Genome 13 (8), 475-478 (2002)

JOURNAL MEDLINE 22213789  
PUBMED 12226715  
COMMENT Contact: Smith TPL  
USDA, ARS, US Meat Animal Research Center  
PO Box 166, Clay Center, NE 68933-0166, USA  
Tel: 402 762 4366  
Fax: 402 762 4390  
Email: smith@mail.marc.usda.gov  
Single pass sequencing. Bases called and alt trimmed with phred  
v0.980904.e. Vector identified by cross\_match with the -mismatch 18  
and -mismatch 12 options.  
PCR Primers  
FORWARD: AGGAACAGCTATGACCAT  
BACKWARD: GTTTCACGACGACGAC  
Plate: 126 row: K column: 16  
Seq primer: ATTAGGTGACACTATAG.  
Location/Qualifiers  
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/clone\_lib="MARC 1P1G"  
/note="Vector: pCMV SPORT6; Site 1: NotI, Site 2: SalI;  
Library made from pooled tissue from day 11, 13, 15, 20,  
and 30 embryos."

## TITLE

JOURNAL MEDLINE 22213789  
PUBMED 12226715  
COMMENT Contact: Smith TPL  
USDA, ARS, US Meat Animal Research Center  
PO Box 166, Clay Center, NE 68933-0166, USA  
Tel: 402 762 4366  
Fax: 402 762 4390  
Email: smith@mail.marc.usda.gov  
Single pass sequencing. Bases called and alt trimmed with phred  
v0.980904.e. Vector identified by cross\_match with the -mismatch 18  
and -mismatch 12 options.  
PCR Primers  
FORWARD: AGGAACAGCTATGACCAT  
BACKWARD: GTTTCACGACGACGAC  
Plate: 126 row: K column: 16  
Seq primer: ATTAGGTGACACTATAG.  
Location/Qualifiers  
1..424  
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/clone\_lib="MARC 1P1G"  
/note="Vector: pCMV SPORT6; Site 1: NotI, Site 2: SalI;  
Library made from pooled tissue from day 11, 13, 15, 20,  
and 30 embryos."

## COMMENT

JOURNAL MEDLINE 22213789  
PUBMED 12226715  
COMMENT Contact: Smith TPL  
USDA, ARS, US Meat Animal Research Center  
PO Box 166, Clay Center, NE 68933-0166, USA  
Tel: 402 762 4366  
Fax: 402 762 4390  
Email: smith@mail.marc.usda.gov  
Single pass sequencing. Bases called and alt trimmed with phred  
v0.980904.e. Vector identified by cross\_match with the -mismatch 18  
and -mismatch 12 options.  
PCR Primers  
FORWARD: AGGAACAGCTATGACCAT  
BACKWARD: GTTTCACGACGACGAC  
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Seq primer: ATTAGGTGACACTATAG.  
Location/Qualifiers  
1..424  
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/clone\_lib="MARC 1P1G"  
/note="Vector: pCMV SPORT6; Site 1: NotI, Site 2: SalI;  
Library made from pooled tissue from day 11, 13, 15, 20,  
and 30 embryos."

## FEATURES

source

## ORIGIN

## Alignment Scores:

Pred. No.: 0.00572 Length: 424  
Score: 13.00 Matches: 13  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 81.25% Indels: 0  
DB: Gaps: 0

US-10-726-967A-3 (1-16) x B1337739 (1-424)

## QY

4 GlyIleArgLeuProLeuArgSerGlyLeuGlyGlyAla 16

DB 376 GGCATCCGGCTGCCCTGCGAAGCGGCTTGGGGGGCA 414

RESULT 4  
LOCUS BG833894 563 bp mRNA linear EST 25-MAY-2001  
DEFINITION BG833894 351953 MARC 1P1G Sus scrofa cDNA 5', mRNA sequence.  
ACCESSION BG833894  
VERSION BG833894.1 GI:14198715  
KEYWORDS EST.  
SOURCE Sus scrofa (pig)  
ORGANISM Sus scrofa

REFERENCE  
AUTHORS Mammalia; Eutheria; Cetartiodactyla; Suina; Suidae; Sus.  
1 (bases 1 to 563)  
Fahnenkrug, S.C., Smith, T.P.L., Freking, B.A., Cho, J., White, J.,  
Vallet, J., Wise, T., Rohrer, G.A., Ferreira, G., Sultana, R.,  
Quackenbush, J., and Keefe, J.W.  
Porcine gene discovery by normalized cDNA-library sequencing and  
EST cluster assembly  
Mamm. Genome 13 (8), 475-478 (2002)

TITLE Mamm. Genome 13 (8), 475-478 (2002)

JOURNAL  
MEDLINE 22213789  
PUBMED 12226715

COMMENT Contact: Smith TPL  
USDA, ARS, US Meat Animal Research Center  
PO Box 166, Clay Center, NE 68933-0166, USA  
Tel: 402 762 4366  
Fax: 402 762 4390  
Email: smith@email.marc.usda.gov  
Single pass sequencing. Bases called and alt trimmed with phred  
v0.980904.e. Vector identified by cross\_match with the -minscore 18  
and -minmatch 12 options.  
PCR Primers  
FORWARD: AGGAACACAGCTATGACCAT  
BACKWARD: GTTTCACGTCACGACG  
Place: 111 row: G column: 20  
Seq primer: ATTAGTGACACTATAG.  
Location/Qualifiers  
source 1..563  
/organism="Sus scrofa"  
/mol\_type="mRNA"  
/db\_xref="taxon:9823"  
/cfeature\_type="pooled"  
/lab\_host="DH10B"  
/clone\_1ib="MARC 1P1G"  
/note="Vector: PCMV SPORT6; Site 1: NotI; Site 2: SalI;  
library made from pooled tissue from day 11, 13, 15, 20,  
and 30 embryos."

ORIGIN  
Alignment Scores:  
Pred. No.: 0.00737 Length: 563  
Score: 13.00 Matches: 13  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 81.25% Indels: 0  
Gaps: 0  
DB: 4

US-10-726-967A-3 (1-16) X BG833894 (1-563)

QY 4 GYIIeArGLEuPProLeuArGSeRGIyLeuGIyGIA1a 16  

DB 504 GGCATCCGGCTGCCCTGCGAAGCGGCTTGGGGGGCA 542

RESULT 5  
LOCUS BP452814 616 bp mRNA linear EST 30-DEC-2003  
DEFINITION BP452814 full-length enriched swine cDNA library, adult liver Sus  
scrofa cDNA clone UMR10122F10 5', mRNA sequence.  
ACCESSION BP452814  
VERSION BP452814.1 GI:40442881  
KEYWORDS EST.  
SOURCE Sus scrofa (pig)  
ORGANISM Sus scrofa

REFERENCE  
AUTHORS Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Cetartiodactyla; Suina; Suidae; Sus.  
1 (bases 1 to 616)  
Uenishi, H., Iguchi, T., Suzuki, K., Sawazaki, T., Toki, D., Shinkai, H.,  
Okumura, N., Hamada, N., and Anata, T.  
PEDS (Pig EST Data Explorer): construction of a database for ESTs  
derived from porcine full-length cDNA libraries  
Nucleic Acids Res. 32 (1), D484-D488 (2004)  
Contact: Hirohide Uenishi  
Animal Genome Laboratory, Genome Research Department  
National Institute of Agrobiological Sciences  
2 Ikenodai, Tsukuba, Ibaraki 305-8602, Japan  
Tel: +81-29-838-8627  
Fax: +81-29-838-8627  
Email: huenishi@affrc.go.jp  
EST project with full-length enriched cDNA libraries carried out in  
Animal Genome Research Program (Japan) by National Institute of  
Agrobiological Sciences and STAFF-Institute  
Single pass sequencing of clones derived from oligo-capped cDNA  
library  
Vector sequences were eliminated by RepeatMasker version 2002/07/13  
and crossmatch version 0.990319  
Low quality bases were trimmed based on the quality values.  
Location/Qualifiers  
source 1..616  
/organism="Sus scrofa"  
/mol\_type="mRNA"  
/db\_xref="taxon:9823"  
/clone\_1ib="UMR10122F10"  
/cfeature\_type="adult"  
/dev\_stage="adult"  
/clone\_1ib="full-length enriched swine cDNA library, adult  
liver"

ORIGIN  
Alignment Scores:  
Pred. No.: 0.00798 Length: 616  
Score: 13.00 Matches: 13  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 81.25% Indels: 0  
Gaps: 5  
DB: 5

US-10-726-967A-3 (1-16) X BP452814 (1-616)

QY 4 GYIIeArGLEuPProLeuArGSeRGIyLeuGIyGIA1a 16  

DB 548 GGCATCCGGCTGCCCTGCGAAGCGGCTTGGGGGGCA 586

RESULT 6  
LOCUS CE200035/c 171 bp DNA linear GSS 25-SEP-2003  
DEFINITION tigr-gss-dog-17000372217391 Dog Library Canis familiaris genomic,  
genomic survey sequence.  
ACCESSION CE200035  
VERSION CE200035.1 GI:35355688  
KEYWORDS GSS.  
SOURCE Canis familiaris (dog)  
ORGANISM Canis familiaris  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Carnivora; Fissipedia; Canidae; Canis.  
1 (bases 1 to 171)  
Kirkness, E.F., Batina, V., Halpern, A.L., Levy, S., Remington, K.,  
Rusch, D.B., Delcher, A.L., Pop, M., Wang, W., Fraser, C.M. and  
Venter, J.C.  
The dog genome: survey sequencing and comparative analysis  
Science 301 (5641), 1898-1903 (2003)

TITLE The dog genome: survey sequencing and comparative analysis  
JOURNAL Science 301 (5641), 1898-1903 (2003)  
MEDLINE 22875432  
PUBMED 14512627

COMMENT Contact: Kirkness EF  
The Institute for Genomic Research  
Department of Eukaryotic Genomics, TIGR, 9712 Medical Center Drive,  
Rockville, MD 20850, USA

Tel: 301-838-0200  
Fax: 301-838-0208  
Email: ekirknes@igr.org  
Class: shotgun.  
Location/Qualifiers  
1. .171

FEATURES  
Source  
/organism="Canis familiaris"  
/mol\_type="genomic DNA"  
/strain="Standard Poodle"  
/db\_xref="taxon:9615"  
/clone\_lib="Dog Library"  
/note="Site 1: BclXI; Libraries were prepared from peripheral blood"

## ALIGNMENT SCORES:

Pred. No.:	0.227	Length:	171
Score:	11.00	Matches:	11
Percent Similarity:	100.00%	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	68.75%	Indels:	0
DB:	9	Gaps:	0

US-10-726-967A-3 (1-16) x CE200035 (1-171)

QY 4 Gly|learg|leup|ro|leu|arg|ser|gly|leu|gly 14  
168 GGCATCGGCTGCCCTCGCGACGCGGCTCGGG 136

RESULT 7  
CN692524 493 bp mRNA linear EST 18-MAY-2004  
LOCUS E0324B10-5 NIA Mouse E10.5 whole embryo cDNA library (Long) Mus  
DEFINITION musculus cDNA clone NIA:E0324B10 IMAGE:30861237 5', mRNA sequence.  
ACCESSION CN692524  
VERSION CN692524.1 GI:47461272  
KEYWORDS EST.  
SOURCE Mus musculus (house mouse)  
ORGANISM Mus musculus  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.

REFERENCE 1 (bases 1 to 493)  
AUTHORS Sharov,A.A., Piao,Y., Matoba,R., Dudekula,D.B., Qian,Y.,  
Vanburen,V., Falco,G., Martin,P.R., Stagg,C.A., Bassez,U.C.,  
Wang,Y., Carter,M.G., Hamatani,T., Alba,K., Akutsu,H., Sharova,L.,  
Tanaka,T.S., Kimber,W.L., Yoshikawa,T., Jaradat,S.A., Pantano,S.,  
Nagaraja,R., Boheler,K.R., Taub,D., Hodess,R.J., Longo,D.L.,  
Schlessinger,D., Keller,J., Klotz,E., Kelsoe,G., Umezawa,A.,  
Vescovi,A.L., Rossant,J., Kunath,T., Hogan,B.L., Curci,A.,  
D'Urso,M., Kelso,J., Hide,W. and Ko,M.S.

TITLE Transcriptional analysis of mouse stem cells and early embryos  
JOURNAL Plos Biol. 1 (3), 410-419 (2003)  
COMMENT Contact: Dawood B. Dudekula  
Laboratory of Genetics  
National Institute on Aging/National Institutes of Health  
333 Cassell Drive, Suite 4000, Baltimore, MD 21224-6820, USA  
Email: cdna@igsn.grc.nia.nih.gov  
Plate: E0324 row: B column: 10  
Seq primer: M13 Reverse  
High quality sequence stop: 493  
POLYA=NO.

FEATURES  
Source  
Location/Qualifiers  
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/organism="Mus musculus"  
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/strain="C57BL/6J"  
/db\_xref="taxon:10090"  
/clone\_lib="NIA:E0324B10 IMAGE:30861237"  
/isue\_type="whole embryo including extramembronic  
tissues at 10.5-days postcoitum"  
/dev\_stage="E10.5"  
/lab\_host="DH10B"

/clone\_lib="NIA Mouse E10.5 whole embryo cDNA library (Long)"  
/note="Vector: pCMV-SPORT6 (Invitrogen), Site 1: SalI; Site 2: NotI, Mouse cDNA project by the Laboratory of Genetics, National Institute on Aging (NIA), Intramural Research Program, NIH (http://igsn.grc.nia.nih.gov/cDNA). This is a long-transcript enriched cDNA library (Ref. Genome Res. 11: 1553-1558 (2001). [PMID: 11544199])". Total RNAs were extracted from a pool of 8 embryos at 10.5-days postcoitum. Double-stranded cDNAs were synthesized with an oligo(dT) primer [Invitrogen: 5'-pACGATGTTCTAGATCGGACGCGCCCTTTTCTTTT-3'] from 2ug of total RNA, treated with T4 DNA polymerase, and purified by ethanol-precipitation. The cDNAs were ligated to lone-linker lp-SalI, purified by phenol/chloroform, and separated from free linkers by Centricon 100. Then, the cDNAs were amplified by long-range high fidelity PCR using Ex Taq polymerase (Takara) with a primer SalI-S. The products were purified by phenol/chloroform and Centricon 100. The cDNAs were digested with SalI and NotI enzymes and cloned into SalI/NotI site of pCMV-SPORT6 plasmid vector. The DH10B E. coli host was transformed with the ligation mixture by the standard chemical method. The average insert size is about 3.4kb. The library was constructed by Yulan Piao."

## ALIGNMENT SCORES:

Pred. No.:	5.47	Length:	493
Score:	10.00	Matches:	10
Percent Similarity:	100.00%	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	62.50%	Indels:	0
DB:	7	Gaps:	0

US-10-726-967A-3 (1-16) x CN692524 (1-493)

QY 4 Gly|learg|leup|ro|leu|arg|ser|gly|leu 13  
382 GGCATCGGCTGCCCTCGCGACGCGCTCG 411

RESULT 8  
CN697484 533 bp mRNA linear EST 18-MAY-2004  
LOCUS E0394H10-5 NIA Mouse E10.5 whole embryo cDNA library (Long) Mus  
DEFINITION musculus cDNA clone NIA:E0394H10 IMAGE:30868029 5', mRNA sequence.  
ACCESSION CN697484  
VERSION CN697484.1 GI:47466233  
KEYWORDS EST.  
SOURCE Mus musculus (house mouse)  
ORGANISM Mus musculus  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.

REFERENCE 1 (bases 1 to 533)  
AUTHORS Sharov,A.A., Piao,Y., Matoba,R., Dudekula,D.B., Qian,Y.,  
Vanburen,V., Falco,G., Martin,P.R., Stagg,C.A., Bassez,U.C.,  
Wang,Y., Carter,M.G., Hamatani,T., Alba,K., Akutsu,H., Sharova,L.,  
Tanaka,T.S., Kimber,W.L., Yoshikawa,T., Jaradat,S.A., Pantano,S.,  
Nagaraja,R., Boheler,K.R., Taub,D., Hodess,R.J., Longo,D.L.,  
Schlessinger,D., Keller,J., Klotz,E., Kelsoe,G., Umezawa,A.,  
Vescovi,A.L., Rossant,J., Kunath,T., Hogan,B.L., Curci,A.,  
D'Urso,M., Kelso,J., Hide,W. and Ko,M.S.

TITLE Transcriptional analysis of mouse stem cells and early embryos  
JOURNAL Plos Biol. 1 (3), 410-419 (2003)  
COMMENT Contact: Dawood B. Dudekula  
Laboratory of Genetics  
National Institute on Aging/National Institutes of Health  
333 Cassell Drive, Suite 4000, Baltimore, MD 21224-6820, USA  
Email: cdna@igsn.grc.nia.nih.gov  
Plate: E0394 row: H column: 10  
Seq primer: M13 Reverse  
High quality sequence stop: 533  
POLYA=NO.

FEATURES  
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Location/Qualifiers  
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/organism="Mus musculus"  
/mol\_type="mRNA"  
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/db\_xref="taxon:10090"  
/db\_xref="taxon:10090"  
/clone="NIA:E0394H10 IMAGE:30868029"  
/tissue\_type="whole embryo including extraembryonic  
tissues at 10.5-days postcoitum"  
/dev\_stage="E10.5"  
/lab\_host="DH10B"  
/clone.lib="NIA Mouse E10.5 whole embryo cDNA library  
(long)"  
/note="Vector: pCMV-SPORT6 (Invitrogen); Site 1: SalI;  
Site 2: NotI; Mouse cDNA project by the Laboratory of  
Genetics, National Institute on Aging (NIA), Intramural  
Research Program, NIH (<http://igsun.grc.nia.nih.gov/cDNA>).  
This is a long-transcript enriched cDNA library (Ref.  
Genome Res. 11: 1553-1558 (2001). [PMD: 11544199]). Total  
RNAs were extracted from a pool of 8 embryos at 10.5-days  
postcoitum. Double-stranded cDNAs were synthesized with an  
Oligo(dT) primer (Invitrogen):  
5'-PACACTGATCTAGATCGAGCGCGCCCTTTT-3' from  
2ug of total RNA, treated with T4 DNA polymerase, and  
purified by ethanol-precipitation. The cDNAs were ligated  
to lone-linker Lr-SalI, purified by phenol/chloroform, and  
separated from free linkers by Centricon 100. Then, the  
cDNAs were amplified by long-range high fidelity PCR using  
Ex Tag polymerase (Takara) with a primer SalI-S. The  
products were purified by phenol/chloroform and Centricon  
100. The cDNAs were digested with SalI and NotI enzymes  
and cloned into SalI/NotI site of pCMV-SPORT6 plasmid  
vector. The DH10B E. coli host was transformed with the  
ligation mixture by the standard chemical method. The  
average insert size is about 3.4kb. The library was  
constructed by Yulan Piao."

ORIGIN  
Alignment Scores:  
Pred. No.: 5.87 Length: 533  
Score: 10.00 Matches: 10  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 62.50% Indels: 0  
DB: 7 Gaps: 0  
US-10-726-967A-3 (1-16) x CN697484 (1-533)

QY 4 GYIIeArlgLeuProLeuArlgSerGlyLeu 13  
|||||  
504 GGCATCGGCTGCTCCCTTCGACGGCGCTG 533

RESULT 9  
CF903755 547 bp mRNA linear EST 04-NOV-2003  
LOCUS A0413D10-5 NIA Mouse Osteoblast cDNA library (Long 1) Mus musculus  
DEFINITION cDNA clone NIA:A0413D10 IMAGE:30739245 5', mRNA sequence.  
ACCESSION CF903755  
VERSION CF903755.1 GI:38170804  
KEYWORDS EST.  
SOURCE Mus musculus (house mouse)  
ORGANISM Mus musculus  
Eukaryota; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
Piao, Y., Ko, N.T., Lim, M.K. and Ko, M.S.H.  
Construction of long-transcript enriched cDNA libraries from  
submicrogram amounts of total RNAs by a universal PCR amplification  
method  
Journal Genome Res. 11 (9), 1553-1558 (2001)  
MEDLINE 21429098  
PUBMED 11544199

COMMENT  
Contact: Dawood B. Dudekula  
Laboratory of Genetics  
National Institute on Aging/National Institutes of Health  
333 Cassell Drive, Suite 4000, Baltimore, MD 21224-6820, USA  
Email: [cdna@igsun.grc.nia.nih.gov](mailto:cdna@igsun.grc.nia.nih.gov)  
Plate: A0413 row: D column: 10  
Seq primer: M13 Reverse  
High quality sequence stop: 547  
POLYA=No.

FEATURES  
source  
Location/Qualifiers  
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/organism="Mus musculus"  
/mol\_type="mRNA"  
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/clone="NIA:A0413D10 IMAGE:30739245"  
/dev\_stage="K05A/A1 cells"  
/lab\_host="DH10B"  
/clone.lib="NIA Mouse Osteoblast cDNA library (long 1)"  
/note="Vector: pCMV-SPORT6 (Invitrogen); Site 1: SalI;  
Site 2: NotI; Mouse cDNA project by the Laboratory of  
Genetics, National Institute on Aging (NIA), Intramural  
Research Program, NIH (<http://igsun.grc.nia.nih.gov/cDNA>).  
This is a long-transcript enriched cDNA library (Ref.  
Genome Res. 11: 1553-1558 (2001). [PMD: 11544199]). Total  
RNAs were obtained from Dr. Akihiro Umezawa (Keio  
University School of Medicine, Japan). Double-stranded  
cDNAs were synthesized with an Oligo(dT) primer  
(Invitrogen):  
5'-PACACTGATCTAGATCGAGCGCGCCCTTTT-3' from  
2.1 ug of total RNA, treated with T4 DNA polymerase, and  
purified by ethanol-precipitation. The cDNAs were ligated  
to lone-linker Lr-SalI, purified by phenol/chloroform, and  
separated from free linkers by Centricon 100. Then, the  
cDNAs were amplified by long-range high fidelity PCR using  
Ex Tag polymerase (Takara) with a primer SalI-S. The  
products were purified by phenol/chloroform and Centricon  
100. The cDNAs were digested with SalI and NotI enzymes  
and cloned into SalI/NotI site of pCMV-SPORT6 plasmid  
vector. The DH10B E. coli host was transformed with the  
ligation mixture by the standard chemical method. The  
average insert size is about 3.0 kb. The library was  
constructed by Yulan Piao."

ORIGIN  
Alignment Scores:  
Pred. No.: 6 Length: 547  
Score: 10.00 Matches: 10  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 62.50% Indels: 0  
DB: 7 Gaps: 0  
US-10-726-967A-3 (1-16) x CF903755 (1-547)

QY 4 GYIIeArlgLeuProLeuArlgSerGlyLeu 13  
|||||  
497 GGCATCGGCTGCTCCCTTCGACGGCGCTG 526

RESULT 10  
CF906581 563 bp mRNA linear EST 04-NOV-2003  
LOCUS A0448H07-5 NIA Mouse Osteoblast cDNA library (Long 1) Mus musculus  
DEFINITION cDNA clone NIA:A0448H07 IMAGE:30742650 5', mRNA sequence.  
ACCESSION CF906581  
VERSION CF906581.1 GI:38173630  
KEYWORDS EST.  
SOURCE Mus musculus (house mouse)  
ORGANISM Mus musculus  
Eukaryota; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
1 (bases 1 to 563)

**AUTHORS** Piao, Y., Ko, N.T., Lim, M.K. and Ko, M.S.H.  
**TITLE** Construction of long-transcript enriched cDNA libraries from submicrogram amounts of total RNAs by a universal PCR amplification method  
**JOURNAL** Genome Res. 11 (9), 1553-1558 (2001)  
**MEDLINE** 21429098  
**PUBMED** 11544199  
**COMMENT** Contact: Dawood B. Dudekula  
 Laboratory of Gene  
 National Institute on Aging/National Institutes of Health  
 333 Cassell Drive, Suite 4000, Baltimore, MD 21224-6820, USA  
 Email: chnag@nigmsun.grc.nia.nih.gov  
 Plate: A0448 Row: H Column: 07  
 Seq primer: M13 Reverse  
 High quality sequence stop: 563  
 POLYA=NO.

**FEATURES**  
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 /organism="Mus musculus"  
 /mol\_type="mRNA"  
 /strain="C3H/He mice"  
 /db\_xref="n1:EST:A0448H07-5"  
 /db\_xref="taxon:10090"  
 /clone="NIA:A0448H07 IMAGE:30742650"  
 /dev\_stage="KUSA/A1 cells"  
 /lab\_host="DH10B"  
 /clone\_lib="NIA Mouse Osteoblast cDNA Library (Long 1)"  
 /note="Vector: PCMV-SPORT6 (Invitrogen); Site 1: SalI; Site 2: NotI; Mouse cDNA project by the Laboratory of Genetics, National Institute on Aging (NIA), Intramural Research Program, NIH (http://igsun.grc.nia.nih.gov/cDNA). This is a long-transcript enriched cDNA library (Ref. Genome Res. 11: 1553-1558 (2001). (PMID: 11544199)). Total RNAs were obtained from Dr. Akihiko Umezawa (Keio University School of Medicine, Japan) Double-stranded cDNAs were synthesized with an Oligo(dt) primer (Invitrogen).  
 5'-pactcctggtctcagatccgagcggccgcccctttttttttttt-3' from 2.1 ug of total RNA, treated with T4 DNA polymerase, and purified by ethanol-precipitation. The cDNAs were ligated to lone-linker IL-SalI, purified by phenol/chloroform, and separated from free linkers by Centricon 100. Then, the cDNAs were amplified by long-range high fidelity PCR using Ex Taq polymerase (Takara) with a primer Sal4-S. The products were purified by phenol/chloroform and Centricon 100. The cDNAs were digested with SalI and NotI enzymes and cloned into SalI/NotI site of PCMV-SPORT6 plasmid vector. The DH10B E. coli host was transformed with the ligation mixture by the standard chemical method. The average insert size is about 3.0 kb. The library was constructed by Yulian Piao."

**ORIGIN**  
 Alignment Scores:  
 Scored. NO.: 6.16 Length: 563  
 Score: 10.00 Matches: 10  
 Percent Similarity: 100.00% Conservative: 0  
 Best Local Similarity: 100.00% Mismatches: 0  
 Query Match: 62.50% Indels: 0  
 Gaps: 7 Gaps: 0

US-10-726-967A-3 (1-16) x CF906581 (1-563)  
 OY 4 GlyTleargleuproleuargserGlyleu 13  
 Db 497 GGCAATCGGCGCTGCCCTTCGACGCGGCTG 526

**RESULT 11** CF171218 576 bp mRNA linear EST 25-JUL-2003  
 LOCUS CF171218  
 DEFINITION B0839F04-5 NIA Mouse Newborn Kidney cDNA library (Long 1) Mus  
 musculus cDNA clone NIA:B0839F04 IMAGE:30471211 5', mRNA sequence.  
 ACCSSION CF171218

VERSION CFI71218.1 GI:33280767  
 KEYWORDS EST.  
 SOURCE Mus musculus (house mouse)  
 ORGANISM Mus musculus  
 Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 Mammalia; Eutheria; Rodentia; Scuriognathi; Muridae; Murinae; Mus.  
 1 (bases 1 to 576)  
 Piao,Y., Ko,N.T., Lim,M.K. and Ko,M.S.H.  
 Construction of long-fragment enriched cDNA libraries from  
 submicrogram amounts of total RNAs by a universal PCR amplification  
 method  
 Genome Res. 11 (9), 1553-1558 (2001)  
 21429098  
 11542199  
 CONTACT Contact: Dawood B. Dudekula  
 LABORATORY Laboratory of Genetics  
 NATIONAL INSTITUTE on Aging/National Institutes of Health  
 333 Cassell Drive, Suite 4000, Baltimore, MD 21224-6820, USA  
 Email: cdna@nigmsun.grc.nia.nih.gov  
 Plate: B0839 Row: F Column: 04  
 Seq primer: M13 Reverse  
 High quality sequence stop: 576  
 POLYVA=No.

FEATURES Location/Qualifiers  
 1..576

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FEATURES
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            /db_xref="taeST:B0839F04-5"
            /db_xref="taxon:10090"
            /clone="NIA:B0839F04 IMAGE:30471231"
            /dev_stage="Newborn Kidney"
            /lab_host="DH10B"
            /clone_lib="NIA Mouse Newborn Kidney cDNA library (long
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            /note="Vector: pCMV-SPORT6 (Invitrogen); Site_1: SalI;
                Site_2: NotI; Mouse cDNA project by the Laboratory of
                Genetics, National Institute on Aging (NIA), Intramural
                Research Program, NIH (http://lgsun.grc.nia.nih.gov/cDNA).
                In brief, double-stranded cDNAs were synthesized with an
                Oligo(dT) primer (Invitrogen):
                5'-GACATGCTTCAGATCGGAGCGGCCGCCCTTTTTTTTTT-3' from
                26 ug of total RNA, treated with T4 DNA polymerase, and
                purified by ethanol-precipitation. The cDNAs were ligated
                to home-linker Lr-Sal4, purified by phenol/chloroform, and
                separated from free linkers by Centricon 100. Then, the
                cDNAs were amplified by long-range high fidelity PCR using
                Ex Taq polymerase (Takara) with a primer Sal4-S. The
                products were purified by phenol/chloroform and Centricon
                100. The cDNAs were digested with SalI and NotI enzymes
                and cloned into SalI/NotI site of pCMV-SPORT6 plasmid
                vector. The DH10B E. coli host was transformed with the
                ligation mixture by the standard chemical method. The
                average insert size is about 3.0 kb. The library was
                constructed by Yulan Piao."

ORIGIN
Alignment Scores:
Pred. NO.:          6.29      Length:       576
Score:              10.00     Matches:      10
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Best local Similarity: 100.00% Mismatches:    0
Query Match:        62.50%    Indels:         0
DB:                  7        Gaps:           0

US-10-726-967A-3 (1-16) x CF171218 (1-576)

QY      4 Gly1LargheupProleuArgsergLyieu 13
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Db      497 GGCAATCCGGCTGCCTTCGACGGGCGCTG 526

RESULT 12
RY713879

```

LOCUS BY713879 600 bp mRNA linear EST 17-DEC-2002  
DEFINITION BY713879 RIKEN full-length enriched, 16 days embryo head Mus  
musculus cDNA clone 4122401C04 5', mRNA sequence.  
ACCESSION BY713879  
VERSION BY713879.1 GI:27126044  
KEYWORDS EST.  
SOURCE Mus musculus (house mouse)  
ORGANISM Mus musculus (house mouse)  
REFERENCE  
AUTHORS  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
1 (bases 1 to 600)  
Okazaki, Y., Furuno, M., Kasukawa, T., Adachi, J., Bono, H., Kondo, S.,  
Nikaido, I., Otsu, N., Saito, R., Suzuki, H., Yamanaka, I.,  
Kiyosawa, H., Yagi, K., Tomaru, Y., Hasegawa, Y., Nogami, A.,  
Schonbach, C., Gojobori, T., Baldarelli, R., Hill, D.P., Bul, C.,  
Hume, D.A., Quackenbush, J., Schriml, L.M., Kanapin, A., Matsuda, H.,  
Bacalov, S., Beisel, K.W., Blake, J.A., Brad, D., Brusci, V.,  
Chochia, C., Corbani, L.E., Cousins, S., Dalla, E., Diegani, T.A.,  
Fletcher, C.F., Forrest, A., Fraser, K.S., Gaasterland, T.,  
Gariboldi, M., Gissi, C., Godzik, A., Gough, J., Grimmond, S.,  
Guelinckich, S., Hirokawa, N., Jackson, I.J., Jarvis, E.D., Kanai, A.,  
Kawai, H., Kawasawa, Y., Kedzierski, R.M., King, B.L., Konagaya, A.,  
Kurochkin, I.V., Lee, Y., Lenhard, B., Lyons, P.A., Maglott, D.R.,  
Mallat, L., Marchionni, L., McKenzie, L., Miki, H., Nagashima, T.,  
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Ravasi, T., Reed, J.C., Reed, D.J., Reid, J., Ring, B.Z., Ringwald, M.,  
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Sultana, R., Takekura, Y., Taylor, M.S., Teasdale, R.D., Tomita, M.,  
Verardo, R., Wagner, L., Wahlestedt, C., Wang, Y., Watanabe, Y.,  
Wells, C., Wilming, L.G., Wynshaw-Boris, A., Yanagisawa, M., Yang, I.,  
Yang, L., Yuan, Z., Zavolan, M., Zhu, Y., Zimmer, A., Carninci, P.,  
Hayatsu, N., Hirozane-Kishikawa, T., Kono, H., Nakamura, M.,  
Sakazume, N., Sato, K., Shiraki, T., Waki, K., Kawai, J., Aizawa, K.,  
Arakawa, T., Fukuda, S., Hara, A., Hashizume, W., Imotani, K., Ishii, Y.,  
Itoh, M., Kagawa, I., Miyazaki, A., Sakai, K., Sasaki, D., Shibata, K.,  
Shinagawa, A., Yasunishi, A., Yoshino, M., Waterston, R., Lander, E.S.,  
Rogers, J., Birney, E. and Hayashizaki, Y.  
Analysis of the mouse transcriptome based on functional annotation  
of 60,770 full-length cDNAs  
Nature 420, 563-573 (2002)  
22354683  
JOURNAL  
MEDLINE  
PUBMED  
TITLE  
COMMENT  
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Tel: 81-45-503-9222  
Fax: 81-45-503-9216  
Email: genome-res@sc.riken.jp, URL: http://genome.gsc.riken.jp/  
Adachi, J., Aizawa, K., Akimura, T., Arakawa, T., Carninci, P.,  
Fukuda, S., Hashizume, W., Hayashida, K., Hirozane, T., Hori, F.,  
Imotani, K., Ishii, Y., Itoh, M., Kagawa, I., Kawai, J., Kojima, Y.,  
Kondo, S., Kono, H., Koya, S., Miyazaki, A., Murata, M., Nakamura, M.,  
Nomura, K., Numazaki, R., Ohno, M., Ohsato, N., Saito, R., Sakazume, N.,  
Sano, H., Sasaki, D., Sato, K., Shibata, K., Shiraki, T., Tagami, M.,  
Takeda, Y., Waki, K., Watanabe, A., Muramatsu, M. and Hayashizaki, Y.  
Direct Submissions  
Computational Analysis of Full-length Mouse cDNAs Compared with  
Human Genome Sequences Mamm. Genome. 12, 673-677 (2001)  
Normalization and subtraction of cap-trapper-selected cDNAs to  
prepare full-length cDNA libraries for rapid discovery of new  
genes. Genome Res. 10 (10), 1617-1630 (2000)  
RIKEN integrated sequence analysis (RISA) system-384-format  
sequencing pipeline with 384 multicapillary sequencer. Genome Res.  
10 (11), 1757-1771 (2000)  
Computer-based methods for the mouse full-length cDNA  
encyclopedia: real-time sequence clustering for construction of a  
nonredundant cDNA library. Genome Res. 11 (2), 281-289 (2001)  
cDNA library was prepared and sequenced in Mouse Genome  
Encyclopedia Project of Genome Exploration Research Group in Riken  
Genomic Sciences Center and Genome Science Laboratory in RIKEN.

FEATURES  
SOURCE  
Location/Qualifiers  
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head"  
/note="Site 1: SalI; Site 2: BamHI. cDNA library was  
prepared and sequenced in Mouse Genome Encyclopedia  
Project of Genome Exploration Research Group in Riken  
Genomic Sciences Center and Genome Science Laboratory in  
RIKEN. Division of Experimental Animal Research in Riken  
contributed to prepare mouse tissues. 1st strand cDNA was  
primed with a primer [5'  
GAGAGAGAGAGATCGATCAAGAGCTCTTTTCTTTTCTTTTNN 3'], cDNA was  
prepared by using trehalose thermo-activated reverse  
transcriptase and subsequently enriched for full-length by  
cap-trapper. Second strand cDNA was prepared with the  
primer adapter of sequence [5'  
GAGAGAGAGATCGATCGATTAATTAATTCATCCCCCCCC 3']. cDNA  
was cloned into the xhoI and BamHI sites. Vector: a  
modified pBluescript KS(+) after bulk excision from lambda  
FLC I"

ORIGIN  
Alignment Scores:  
Pred. No.: 6.52 Length: 600  
Score: 10.00 Matches: 10  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 62.50% Indels: 0  
DB: Gaps: 0  
US-10-726-967A-3 (1-16) x BY713879 (1-600)  
QY 4 Gly11eAtgLeuProLeuArgSerGlyLeu 13  
DB 498 GGCATCCGCGCTGCCCTTCGACGCGCTG 527  
RESULT 13  
BB644736  
LOCUS BB644736 619 bp mRNA linear EST 26-OCT-2001  
DEFINITION BB644736 RIKEN full-length enriched, adult male corpora  
quadrigemina Mus musculus cDNA clone B230346M13 5', mRNA sequence.  
ACCESSION BB644736  
VERSION BB644736.1 GI:16479273  
KEYWORDS EST.  
SOURCE Mus musculus (house mouse)  
ORGANISM Mus musculus (house mouse)  
REFERENCE  
AUTHORS  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
1 (bases 1 to 619)  
Arakawa, T., Carninci, P., Fukuda, S., Furuno, M., Hanagaki, T.,  
Hara, A., Hiramoto, K., Hori, F., Ishii, Y., Ito, M., Kawai, J.,  
Kono, H., Konda, M., Koya, S., Matsuyama, T., Miyazaki, A., Nomura, K.,  
Ohno, M., Okazaki, Y., Okido, T., Saito, R., Sakai, C., Sakai, K.,  
Sano, H., Sasaki, D., Shibata, K., Shinagawa, A., Shiraki, T.,  
Sogabe, Y., Suzuki, H., Tagami, M., Tagawa, A., Takahashi, F.,  
Takeda, Y., Tanaka, T., Toyata, T., Muramatsu, M. and Hayashizaki, Y.  
RIKEN Mouse ESTs (Arakawa, T., et al. 2001)  
Unpublished (2001)  
TITLE  
JOURNAL  
COMMENT  
Contact: Yoshihide Hayashizaki  
Laboratory for Genome Exploration Research Group, RIKEN Genomic







ORIGIN

GAGAGAGAGAGATCCAGAAGCTCTTTTCTTTTCTTTTNN 3', cDNA was prepared by using trehalose thermo-activated reverse transcriptase and subsequently enriched for full-length by cap-trapper. cDNA went through one round of normalization to Rot = 20.0 and subtraction to Rot = 459.0. Second strand cDNA was prepared with the primer adapter of sequence [5' GAGAGAGAGATCTCCAGTTAATTAATTAATCCCCCCCCCCCC 3']. cDNA was cleaved with XhoI and BamHI. Vector: a modified pBluescript KS(+) after bulk excision from Lambda

FLC I. "

Alignment Scores:

Pred. No.:	6.89	Length:	639
Score:	10.00	Matches:	10
Percent Similarity:	100.00%	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	62.50%	Indels:	0
DB:	2	Gaps:	0

US-10-726-967A-3 (1-16) x BB632244 (1-639)

QY 4 G1Y1leargleupProLeuAArgserGlyLeu 13  
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518 GGCATCCGGCTGCCCTTCGCAGCGGCCTG 547

Search completed: July 27, 2005, 20:35:53

Job time : 3150 secs

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GenCore version 5.1.6  
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OM protein - nucleic search, using frame\_plus.p2n model

Run on: July 27, 2005, 16:41:37 ; Search time 1935 Seconds

(without alignments)  
701.161 Million cell updates/sec

Title: US-10-726-967A-52

Perfect score: 144  
Sequence: 1 GYVEMTVGSPQPTINILVDTCSSNPV 28

Scoring table:  
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Xgapop 10.0 , Xgapext 0.5  
Ygapop 10.0 , Ygapext 0.5  
Fgapop 6.0 , Fgapext 7.0  
Delop 6.0 , Delext 7.0

Searched: 4708233 seqs, 24227607955 residues  
Total number of hits satisfying chosen parameters: 9416466

Minimum DB seq length: 0  
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Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Command line parameters:  
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-Q/cg2.1/USPTO.spool.p/US10726967/unal.26072005.130734.6141/app.query.fasta.1.199  
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Database :

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7: gb\_ph:\*  
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13: gb\_un:\*  
14: gb\_vi:\*

Prod. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

# SUMMARIES

Result No.	Score	Query Match	Length	ID	Description
1	144	100.0	517	9	AB089958 Homo sapi
2	144	100.0	1278	6	BD235898 Alzheimer
3	144	100.0	1278	6	AR224104 Sequence
4	144	100.0	1278	6	AR269235 Sequence

5	144	100.0	1278	6	AR478790	Sequence
6	144	100.0	1278	6	AR487356	Sequence
7	144	100.0	1278	6	AR531996	Sequence
8	144	100.0	1278	6	AR540897	Sequence
9	144	100.0	1278	6	AR560107	Sequence
10	144	100.0	1278	6	AX105409	Sequence
11	144	100.0	1278	6	AX573847	Sequence
12	144	100.0	1287	6	AR224122	Sequence
13	144	100.0	1287	6	AR269253	Sequence
14	144	100.0	1287	6	AR478808	Sequence
15	144	100.0	1287	6	AR487374	Sequence
16	144	100.0	1287	6	AR532014	Sequence
17	144	100.0	1287	6	AR540915	Sequence
18	144	100.0	1287	6	AR560125	Sequence
19	144	100.0	1287	6	AX105432	Sequence
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23	144	100.0	1302	6	AR269234	Sequence
24	144	100.0	1302	6	AR478789	Sequence
25	144	100.0	1302	6	AR487355	Sequence
26	144	100.0	1302	6	AR531995	Sequence
27	144	100.0	1302	6	AR540896	Sequence
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36	144	100.0	1305	6	AR532015	Sequence
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38	144	100.0	1305	6	AR560126	Sequence
39	144	100.0	1305	6	AX105434	Sequence
40	144	100.0	1305	6	AX573872	Sequence
41	144	100.0	1333	9	AB050438	Homo sapi
42	144	100.0	1341	6	BD235895	Alzheimer
43	144	100.0	1341	6	AR224101	Sequence
44	144	100.0	1341	6	AR269232	Sequence
45	144	100.0	1341	6	AR478787	Sequence

## ALIGNMENTS

RESULT 1	AB089958	517 bp	mRNA	linear	PRI 19-AUG-2003
LOCUS	AB089958				
DEFINITION	Homo sapiens BACE mRNA for beta-site APP cleaving enzyme isoform I-127, complete cds.				
ACCESSION	AB089958				
VERSION	AB089958.1	GI:34014375			
KEYWORDS	Homo sapiens (human)				
SOURCE	Homo sapiens				
ORGANISM	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Primates; Catarrhini; Hominiidae; Homo.				
REFERENCE	1 Tanahashi, H.				
AUTHORS	A novel alternatively spliced isoform of BACE, I-127 induced by cycloheximide treatment				
TITLE	Unpublished				
JOURNAL	2 (bases 1 to 517)				
REFERENCE	Tanahashi, H.				
AUTHORS	Submitted (17-AUG-2002) Hiroshi Tanahashi, National Institute of Neuroscience, Division of Demyelinating Disease and Aging; 4-1-1 Ogawahigashi, Kodaira, Tokyo 187-8502, Japan				
JOURNAL	(E-mail:tanahashencnp.go.jp, Tel:81-042-341-2711(ex.5163), Fax:81-042-346-1747)				
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ORIGIN

Alignment Scores:  
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Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 9 Gaps: 0

US-10-726-967A-52 (1-28) x AB089958 (1-517)

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QY 21 ThrglySerSerAaspheAlaVal 28  
Db 280 ACAGGACAGCACTTTCAGCTG 303

RESULT 2  
BD235898  
LOCUS BD235898 1278 bp DNA linear PAT 17-JUL-2003  
DEFINITION Alzheimer's disease secretase.  
ACCESSION BD235898.1 GI:33045668  
VERSION BD235898.1  
KEYWORDS JP 2002526081-A/14.  
SOURCE Homo sapiens (human)  
ORGANISM Homo sapiens  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.  
Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and  
Yan,R.  
Alzheimer's disease secretase  
Patent: JP 2002526081-A 14 20-AUG-2002;  
PHARMACIA AND UPJOHN CO  
OS Homo sapiens (human)  
PN JP 2002526081-A/14  
PD 20-AUG-2002  
PR 23-SEP-1999 JP 2000574268  
PF 24-SEP-1998 US 60/101594  
PI MARK E GURNEY, MICHAEL JEROME BIENKOWSKI, ROBERT LEROY PI  
HEINRICHSON,  
PI LUIS A PARODI, RIGIANG YAN  
PC C12N15/09, A61K45/00, A61P25/28, C07K14/47, C07K16/18, C12N1/15, PC  
C12N1/19,  
PC C12N1/21, C12N5/10, C12N9/64, C12P21/02, C12P21/08, C12Q1/37, G01N33/ PC  
15,  
PC G01N33/50// (C12N1/21, C12P1/19), C12N5/00, C12N5/00 CC  
Alzheimer's disease secretase  
FH Key Location/Qualifiers  
FT source 1..1278

FEATURES FT Location/Qualifiers  
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ORIGIN

Alignment Scores:  
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Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
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US-10-726-967A-52 (1-28) x AR224104 (1-1278)

QY 1 G|Y|T|Y|T|Y|V|A|G|U|N|E|T|H|V|A|G|I|S|E|P|P|R|O|G|I|N|T|H|L|E|U|A|N|I|L|E|U|V|A|L|A|P 20  
Db 136 GGGTACTACTAGTGGAGATGACCGTGGCGACCCCGCAGACGCTCAACATCCTGTGGAT 195

QY 21 ThrglySerSerAaspheAlaVal 28  
Db 196 ACAGGACAGCACTTTCAGCTG 219

RESULT 4  
AR269235  
LOCUS AR269235 1278 bp DNA linear PAT 10-APR-2003  
DEFINITION Sequence 27 from patent US 6500667.  
ACCESSION AR269235  
VERSION AR269235.1 GI:29700203  
KEYWORDS

ORIGIN

Alignment Scores:  
Pred. No.: 3.18e-14 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-52 (1-28) x AR224104 (1-1278)

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Db 136 GGGTACTACTAGTGGAGATGACCGTGGCGACCCCGCAGACGCTCAACATCCTGTGGAT 195

QY 21 ThrglySerSerAaspheAlaVal 28  
Db 196 ACAGGACAGCACTTTCAGCTG 219

RESULT 4  
AR269235  
LOCUS AR269235 1278 bp DNA linear PAT 10-APR-2003  
DEFINITION Sequence 27 from patent US 6500667.  
ACCESSION AR269235  
VERSION AR269235.1 GI:29700203  
KEYWORDS

SOURCE Unknown.  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1278)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
TITLE Aspartyl protease 2 (Asp2) antisense oligonucleotides  
JOURNAL Patent: US 6500667-A 27 31-DEC-2002;  
FEATURES Location/Qualifiers  
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ORIGIN

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Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-52 (1-28) x AR269235 (1-1278)

QY 1 G|Y|T|Y|T|V|A|G|U|N|E|T|H|V|A|G|I|S|E|R|P|R|O|G|I|N|T|H|L|E|U|A|N|I|L|E|U|V|A|A|P 20  
DB 136 G|G|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|G|G|G|C|A|G|C|C|C|C|C|C|C|C|A|G|A|C|G|T|C|A|A|C|A|T|C|C|T|G|G|T|G|A|T 195

QY 21 T|H|G|I|S|E|R|S|E|R|A|N|P|H|E|A|V|A| 28  
DB 196 A|C|A|G|G|C|A|G|C|A|G|T|T|T|G|C|A|G|T|G 219

RESULT 5  
AR478790  
LOCUS AR478790 1278 bp DNA linear PAT 14-MAY-2004  
DEFINITION Sequence 27 from patent US 669671.  
ACCESSION AR478790  
VERSION AR478790.1 GI:47237510  
KEYWORDS  
SOURCE Unknown.  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1278)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
TITLE Alzheimer's disease secretase, APP substrates therefor, and uses therefor  
JOURNAL Patent: US 669671-A 27 02-MAR-2004;  
FEATURES Location/Qualifiers  
source 1..1278  
/organism="unknown"  
/mol\_type="genomic DNA"

ORIGIN

Alignment Scores:  
Pred. No.: 3,18e-14 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-52 (1-28) x AR478790 (1-1278)

QY 1 G|Y|T|Y|T|V|A|G|U|N|E|T|H|V|A|G|I|S|E|R|P|R|O|G|I|N|T|H|L|E|U|A|N|I|L|E|U|V|A|A|P 20  
DB 136 G|G|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|G|G|G|C|A|G|C|C|C|C|C|C|C|C|A|G|A|C|G|T|C|A|A|C|A|T|C|C|T|G|G|T|G|A|T 195

QY 21 T|H|G|I|S|E|R|S|E|R|A|N|P|H|E|A|V|A| 28  
DB 196 A|C|A|G|G|C|A|G|C|A|G|T|T|T|G|C|A|G|T|G 219

RESULT 6

AR487356  
LOCUS AR487356 1278 bp DNA linear PAT 14-MAY-2004  
DEFINITION Sequence 27 from patent US 6706485.  
ACCESSION AR487356  
VERSION AR487356.1 GI:47252454  
KEYWORDS  
SOURCE Unknown.  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1278)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
TITLE Method of identifying agents that inhibit APP processing activity  
JOURNAL Patent: US 6706485-A 27 16-MAR-2004;  
FEATURES Location/Qualifiers  
source 1..1278  
/organism="unknown"  
/mol\_type="genomic DNA"

ORIGIN

Alignment Scores:  
Pred. No.: 3,18e-14 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-52 (1-28) x AR487356 (1-1278)

QY 1 G|Y|T|Y|T|V|A|G|U|N|E|T|H|V|A|G|I|S|E|R|P|R|O|G|I|N|T|H|L|E|U|A|N|I|L|E|U|V|A|A|P 20  
DB 136 G|G|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|G|G|G|C|A|G|C|C|C|C|C|C|C|C|A|G|A|C|G|T|C|A|A|C|A|T|C|C|T|G|G|T|G|A|T 195

QY 21 T|H|G|I|S|E|R|S|E|R|A|N|P|H|E|A|V|A| 28  
DB 196 A|C|A|G|G|C|A|G|C|A|G|T|T|T|G|C|A|G|T|G 219

RESULT 7  
AR531996  
LOCUS AR531996 1278 bp DNA linear PAT 08-OCT-2004  
DEFINITION Sequence 27 from patent US 6727074.  
ACCESSION AR531996  
VERSION AR531996.1 GI:53920530  
KEYWORDS  
SOURCE Unknown.  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1278)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
TITLE Alzheimer's disease secretase, APP substrates therefor, and uses therefor  
JOURNAL Patent: US 6727074-A 27 27-APR-2004;  
FEATURES Location/Qualifiers  
source 1..1278  
/organism="unknown"  
/mol\_type="genomic DNA"

ORIGIN

Alignment Scores:  
Pred. No.: 3,18e-14 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-52 (1-28) x AR531996 (1-1278)

QY 1 G|Y|T|Y|T|V|A|G|U|N|E|T|H|V|A|G|I|S|E|R|P|R|O|G|I|N|T|H|L|E|U|A|N|I|L|E|U|V|A|A|P 20  
DB 136 G|G|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|G|G|G|C|A|G|C|C|C|C|C|C|C|C|A|G|A|C|G|T|C|A|A|C|A|T|C|C|T|G|G|T|G|A|T 195

Qy 21 ThrGlySerSerAanPheAlaVal 28  
Db 196 ACAGGACGACGTAACCTTGCACTG 219

RESULT 8  
ARS40897  
LOCUS ARS40897 1278 bp DNA linear PAT 08-OCT-2004  
DEFINITION Sequence 27 from patent US 6737510.  
ACCESSION ARS40897  
VERSION ARS40897.1 GI:53932410  
KEYWORDS  
SOURCE  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1278)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
TITLE Alzheimer's disease secretase, APP substrates therefor, and uses thereof  
JOURNAL Patent: US 6737510-A 27 18-MAY-2004;  
FEATURES  
source  
/organism="unknown"  
/mol\_type="genomic DNA"

ORIGIN  
Alignment Scores:  
Pred. No.: 3,18e-14 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-52 (1-28) x ARS40897 (1-1278)

Qy 1 GlyTyrTyrValGluMetThrValGlySerProProGlnThrLeuAanlleuValAsp 20  
Db 136 GGCTACTACGTGAGATGACCGTGGGACCCCCCGACAGCCTCAACATCTGTTGGAT 195

Qy 21 ThrGlySerSerAanPheAlaVal 28  
Db 196 ACAGGACGACGTAACCTTGCACTG 219

RESULT 9  
ARS60107  
LOCUS ARS60107 1278 bp DNA linear PAT 08-OCT-2004  
DEFINITION Sequence 27 from patent US 6753163.  
ACCESSION ARS60107  
VERSION ARS60107.1 GI:53970474  
KEYWORDS  
SOURCE  
ORGANISM Unknown.  
REFERENCE 1 (bases 1 to 1278)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
TITLE Alzheimer's disease secretase, APP substrates therefor, and uses thereof  
JOURNAL Patent: US 6753163-A 27 22-JUN-2004;  
FEATURES  
source  
/organism="unknown"  
/mol\_type="genomic DNA"

ORIGIN  
Alignment Scores:  
Pred. No.: 3,18e-14 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 0 Gaps: 0

DB: 6 Gaps: 0

US-10-726-967A-52 (1-28) x ARS60107 (1-1278)

Qy 1 GlyTyrTyrValGluMetThrValGlySerProProGlnThrLeuAanlleuValAsp 20  
Db 136 GGCTACTACGTGAGATGACCGTGGGACCCCCCGACAGCCTCAACATCTGTTGGAT 195

Qy 21 ThrGlySerSerAanPheAlaVal 28  
Db 196 ACAGGACGACGTAACCTTGCACTG 219

RESULT 10  
AX105409  
LOCUS AX105409 1278 bp DNA linear PAT 30-APR-2001  
DEFINITION Sequence 27 from Patent WO0123533.  
ACCESSION AX105409  
VERSION AX105409.1 GI:13921523  
KEYWORDS  
SOURCE  
ORGANISM Homo sapiens (human)  
REFERENCE 1  
AUTHORS Gurney,M. and Bienkowski,M.J.  
TITLE Alzheimer's disease secretase, app substrates therefor, and uses thereof  
JOURNAL Patent: WO 0123533-A 27 05-APR-2001;  
FEATURES  
source  
/organism="Homo sapiens"  
/mol\_type="unassigned DNA"  
/db\_xref="taxon:9606"

ORIGIN  
Alignment Scores:  
Pred. No.: 3,18e-14 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967A-52 (1-28) x AX105409 (1-1278)

Qy 1 GlyTyrTyrValGluMetThrValGlySerProProGlnThrLeuAanlleuValAsp 20  
Db 136 GGCTACTACGTGAGATGACCGTGGGACCCCCCGACAGCCTCAACATCTGTTGGAT 195

Qy 21 ThrGlySerSerAanPheAlaVal 28  
Db 196 ACAGGACGACGTAACCTTGCACTG 219

RESULT 11  
AX573847  
LOCUS AX573847 1278 bp DNA linear PAT 07-JAN-2003  
DEFINITION Sequence 27 from Patent EP1249498.  
ACCESSION AX573847  
VERSION AX573847.1 GI:27551489  
KEYWORDS  
SOURCE  
ORGANISM Homo sapiens (human)  
REFERENCE 1  
AUTHORS Gurney,M. and Bienkowski,M.J.  
TITLE Alzheimer's disease secretase, app substrates therefor, and uses thereof  
JOURNAL Patent: EP 1249498-A 27 16-OCT-2002;  
FEATURES  
source  
/organism="Homo sapiens"  
/mol\_type="unassigned DNA"  
/db\_xref="taxon:9606"

ORIGIN  
/organism="Homo sapiens"  
/mol\_type="unassigned DNA"  
/db\_xref="taxon:9606"

Alignment Scores:  
Pred. No.: 3.18e-14 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967a-52 (1-28) x AX573847 (1-1278)

QY 1 G|Y|T|T|Y|V|A|G|U|W|E|T|H|R|V|A|G|Y|S|E|R|P|R|O|G|I|N|T|H|R|E|U|S|N|I|L|E|U|V|A|A|P 20  
DB 136 GGCCTACTACGTGAGATGACCGTGGCGACCCCGCCGACAGCCTCAACATCTCTGTGAT 195

QY 21 ThrGlySerSerAspAspAlaVal 28  
DB 196 ACAGGACGACGTAACTTTGCAGTG 219

RESULT 12  
LOCUS AR224122 1287 bp DNA linear PAT 26-SEP-2002  
DEFINITION Sequence 50 from patent US 6440698.  
ACCESSION AR224122  
VERSION AR224122.1 GI:23332782  
KEYWORDS  
SOURCE Unknown.  
ORGANISM Unclassified.

REFERENCE 1 (bases 1 to 1287)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.

TITLE Alzheimer's disease secretase, APP substrates therefor, and uses

JOURNAL Patent: US 6440698-A 50 27-AUG-2002;

FEATURES  
source Location/Qualifiers  
1..1287  
/organism="unknown"  
/mol\_type="genomic DNA"

# ORIGIN

Alignment Scores:  
Pred. No.: 3.2e-14 Length: 1287  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967a-52 (1-28) x AR224122 (1-1287)

QY 1 G|Y|T|T|Y|V|A|G|U|W|E|T|H|R|V|A|G|Y|S|E|R|P|R|O|G|I|N|T|H|R|E|U|S|N|I|L|E|U|V|A|A|P 20  
DB 220 GGCCTACTACGTGAGATGACCGTGGCGACCCCGCCGACAGCCTCAACATCTCTGTGAT 279

QY 21 ThrGlySerSerAspAspAlaVal 28  
DB 280 ACAGGACGACGTAACTTTGCAGTG 303

RESULT 13  
LOCUS AR269253 1287 bp DNA linear PAT 10-APR-2003  
DEFINITION Sequence 50 from patent US 650667.  
ACCESSION AR269253  
VERSION AR269253.1 GI:29700221  
KEYWORDS  
SOURCE Unknown.  
ORGANISM Unclassified.

REFERENCE 1 (bases 1 to 1287)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
TITLE Aspartyl protease 2 (Asp2) antisense oligonucleotides  
JOURNAL Patent: US 650667-A 50 31-DEC-2002;  
FEATURES  
source Location/Qualifiers  
1..1287  
/organism="unknown"  
/mol\_type="genomic DNA"

# ORIGIN

Alignment Scores:  
Pred. No.: 3.2e-14 Length: 1287  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967a-52 (1-28) x AR269253 (1-1287)

QY 1 G|Y|T|T|Y|V|A|G|U|W|E|T|H|R|V|A|G|Y|S|E|R|P|R|O|G|I|N|T|H|R|E|U|S|N|I|L|E|U|V|A|A|P 20  
DB 220 GGCCTACTACGTGAGATGACCGTGGCGACCCCGCCGACAGCCTCAACATCTCTGTGAT 279

QY 21 ThrGlySerSerAspAspAlaVal 28  
DB 280 ACAGGACGACGTAACTTTGCAGTG 303

RESULT 14  
LOCUS AR478808 1287 bp DNA linear PAT 14-MAY-2004  
DEFINITION Sequence 50 from patent US 669671.  
ACCESSION AR478808  
VERSION AR478808.1 GI:47237528  
KEYWORDS  
SOURCE Unknown.  
ORGANISM Unclassified.

REFERENCE 1 (bases 1 to 1287)  
AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.

TITLE Alzheimer's disease secretase, APP substrates therefor, and uses

JOURNAL Patent: US 669671-A 50 02-MAR-2004;

FEATURES  
source Location/Qualifiers  
1..1287  
/organism="unknown"  
/mol\_type="genomic DNA"

# ORIGIN

Alignment Scores:  
Pred. No.: 3.2e-14 Length: 1287  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 6 Gaps: 0

US-10-726-967a-52 (1-28) x AR478808 (1-1287)

QY 1 G|Y|T|T|Y|V|A|G|U|W|E|T|H|R|V|A|G|Y|S|E|R|P|R|O|G|I|N|T|H|R|E|U|S|N|I|L|E|U|V|A|A|P 20  
DB 220 GGCCTACTACGTGAGATGACCGTGGCGACCCCGCCGACAGCCTCAACATCTCTGTGAT 279

QY 21 ThrGlySerSerAspAspAlaVal 28  
DB 280 ACAGGACGACGTAACTTTGCAGTG 303

RESULT 15  
LOCUS AR487374 1287 bp DNA linear PAT 14-MAY-2004  
DEFINITION Sequence 50 from patent US 6706485.

ACCESSION AR487374  
 VERSION AR487374.1 GI:47252472  
 KEYWORDS  
 SOURCE  
 ORGANISM Unknown.  
 REFERENCE 1 (bases 1 to 1287)  
 AUTHORS Gurney,M.E., Bienkowski,M.J., Heinrichson,R.L., Parodi,L.A. and Yan,R.  
 TITLE Method of identifying agents that inhibit APP processing activity  
 JOURNAL Patent: US 6706485-A 50 16-MAR-2004;  
 FEATURES Location/Qualifiers  
 source 1..1287  
 /organism="Unknown"  
 /mol\_type="genomic DNA"

ORIGIN  
 Alignment Scores:  
 Pred. No.: 3.2e-14 Length: 1287  
 Score: 144.00 Matches: 28  
 Percent Similarity: 100.00% Conservative: 0  
 Best Local Similarity: 100.00% Mismatches: 0  
 Query Match: 100.00% Indels: 0  
 Gaps: 0  
 DB: 6

US-10-726-967a-52 (1-28) x AR487374 (1-1287)

QY 1 GlyTyrTyrValGluMetThrValGlySerProProGlnThrLeuAsnIleuValAsp 20  
 |||||  
 DB 220 GGCTACTAGTGAAGATGACCGTGGGACAGCCCCCGCAGACGCTCAACATCTGTGGAT 279  
 QY 21 ThrGlySerSerSerAsnPheAlaVal 28  
 |||||  
 DB 280 ACAGGACGAGTACTTTCAGATG 303

Search completed: July 27, 2005, 18:50:18  
 Job time : 1939 secs



GenCore version 5.1.6  
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OM protein - nucleic search, using frame\_plus.p2n model

Run on: July 27, 2005, 12:14:54 ; Search time 437 Seconds

(without alignments)  
379.297 Million cell updates/sec

Title: US-10-726-967A-52

Sequence: 1 GYVEMTGSPPQTINILVDTCSSNFAV 28

Scoring table:

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Ygapop 10.0	Ygapext 0.5	
Fgapop 6.0	Fgapext 7.0	
Delop 6.0	Delext 7.0	

Searched: 4390206 seqs, 2959870667 residues

Total number of hits satisfying chosen parameters: 8780412

Minimum DB seq length: 0

Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

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-Q/cg2.1/USPTO.spool.p/US10726567/funat.26072005.130733.6129/app.query.fasta.1.199  
-DB=N.Geneseg.16Dec04 -QFMT=fasta -SUPFIX=p2n.rng -MINMATCH=0.1 -IOOPT=0  
-IOPEXT=0 -INITS=bits -START=1 -END=1 -MATRIX=biolum62 -TRANS=human4.0.cdi  
-LIST=45 -DOCALLIGN=200 -THR\_SCORE=pct -THR\_MAX=100 -THR\_MIN=0 -ALIGN=15  
-MODE=LOCAL -OUTFMT=pct -NORM=ext -HEAPSIZE=500 -MINLEN=0 -MAXLEN=200000000  
-USER=US10726567 @CGN 1.1.708 @runat.26072005.130733.6129 -NCPU=6 -ICPU=3  
-NO MAP -LARGEOUTERY -NEG\_SCORES=0 -WAIT -DSPBLOCK=100 -LONGLOG  
-DEV TIMEOUT=120 -WARN TIMEOUT=30 -THREADS=1 -XGAPOP=10 -XGAPEXT=0.5 -FGAPOP=6  
-FAPEXT=7 -GGAPOP=10 -YGAPEXT=0.5 -DELOP=6 -DELEXT=7

Database : N.Geneseg.16Dec04:\*

1: geneseg1980s:\*\n2: geneseg1990s:\*\n3: geneseg2000s:\*\n4: geneseg2001s:\*\n5: geneseg2001bs:\*\n6: geneseg2002as:\*\n7: geneseg2002bs:\*\n8: geneseg2003as:\*\n9: geneseg2003bs:\*\n10: geneseg2003cs:\*\n11: geneseg2003ds:\*\n12: geneseg2004as:\*\n13: geneseg2004bs:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

# SUMMARIES

Result No.	Score	Query Match	Length	DB	ID	Description
1	144	100.0	1278	3	AAAI5677	AAAI5677 Human Asp
2	144	100.0	1278	4	AAAI1714	AAAI1714 DNA encod
3	144	100.0	1278	4	AAAI1787	AAAI1787 T7-Caspa
4	144	100.0	1278	4	AAAI3033	AAAI3033 T7-Caspa
5	144	100.0	1278	4	AAAI6751	AAAI6751 T7-Caspa

6	144	100.0	1278	4	AAAI529	AAAI529 T7-Caspa
7	144	100.0	1278	6	ABLS2469	ABLS2469 T7-Caspa
8	144	100.0	1278	12	ADJ94339	ADJ94339 Human T7-
9	144	100.0	1278	12	ADOS0435	ADOS0435 T7-Caspa
10	144	100.0	1278	13	ADR75348	ADR75348 T7-Caspa
11	144	100.0	1287	4	AAAI7895	AAAI7895 Human-Asp
12	144	100.0	1287	4	AAAI3276	AAAI3276 Human-Asp
13	144	100.0	1287	4	AAAI6768	AAAI6768 Human-Asp
14	144	100.0	1287	4	AAAI1547	AAAI1547 Human-Asp
15	144	100.0	1287	6	ABLS2487	ABLS2487 Human-Asp
16	144	100.0	1287	12	ADJ94362	ADJ94362 Human-Asp
17	144	100.0	1287	12	ADOS0458	ADOS0458 Human-Asp
18	144	100.0	1287	13	ADR75371	ADR75371 Human-Asp
19	144	100.0	1302	3	AAAI5670	AAAI5670 Human-Asp
20	144	100.0	1302	4	AAAI1773	AAAI1773 DNA encod
21	144	100.0	1302	4	AAAI7876	AAAI7876 Human-pro
22	144	100.0	1302	4	AAAI3032	AAAI3032 Human-pro
23	144	100.0	1302	4	AAAI6750	AAAI6750 Human-pro
24	144	100.0	1302	4	AAAI1528	AAAI1528 Human-CDN
25	144	100.0	1302	6	ABLS2468	ABLS2468 Human-pro
26	144	100.0	1302	12	ADJ94337	ADJ94337 Human-pro
27	144	100.0	1302	12	ADOS0433	ADOS0433 Human-pro
28	144	100.0	1302	13	ADR75346	ADR75346 Human-pro
29	144	100.0	1305	4	AAAI1733	AAAI1733 DNA encod
30	144	100.0	1305	4	AAAI7896	AAAI7896 Human-Asp
31	144	100.0	1305	4	AAAI3277	AAAI3277 Human-Asp
32	144	100.0	1305	4	AAAI6769	AAAI6769 Human-Asp
33	144	100.0	1305	4	AAAI1548	AAAI1548 Human-CDN
34	144	100.0	1305	6	ABLS2468	ABLS2468 Human-Asp
35	144	100.0	1305	12	ADJ94364	ADJ94364 Human-pro
36	144	100.0	1305	12	ADOS0460	ADOS0460 Human-Asp
37	144	100.0	1305	13	ADR75373	ADR75373 Human-Asp
38	144	100.0	1341	3	AAAI5668	AAAI5668 T7-Caspa
39	144	100.0	1341	4	AAAI1711	AAAI1711 DNA encod
40	144	100.0	1341	4	AAAI7874	AAAI7874 T7-Human-
41	144	100.0	1341	4	AAAI3030	AAAI3030 T7-Human-
42	144	100.0	1341	4	AAAI6748	AAAI6748 T7-Human-
43	144	100.0	1341	4	AAAI1526	AAAI1526 Human-CDN
44	144	100.0	1341	6	ABLS2466	ABLS2466 T7-human-
45	144	100.0	1341	12	ADJ94333	ADJ94333 Human-CDN

## ALIGNMENTS

RESULT 1	AAAI5677	standard; DNA; 1278 BP.
ID	AAAI5677	
XX	AAAI5677	
AC	AAAI5677	
DT	03-AUG-2000	(first entry)
XX		
DE	Human Asp2 nucleotide sequence containing proteolytic cleavage site.	
XX		
KW	Aspartyl protease; aspartase; amyloid precursor protein; APP; Asp 2;	
KW	Alzheimer's disease; beta secretase site; ss.	
XX		
OS	Homo sapiens.	
XX		
PN	MO200017369-A2.	
XX		
PD	30-MAR-2000.	
XX		
PF	23-SEP-1999;	99MO-US020881.
XX		
PR	24-SEP-1998;	98US-0101594P.
XX		
PA	(PHAA ) PHARMACIA & UPJOHN CO.	
XX		
PI	Gurney ME, Bienkowski MJ, Heintzson RL, Parodi LA, Yan R;	
DR	WPI; 2000-303209/26.	
DR	P-PSDB; AAY88437.	

XX New enzyme designated human aspartase useful in research into Alzheimer's  
PT Disease is capable of cleaving amyloid protein precursor at the beta  
PT secretase site to produce amyloid beta peptide.  
XX

Example 9; Page 165; 183pp; English.

XX This sequence represents a modified version of the human aspartase 2  
CC (Asp2) nucleotide sequence. The sequence is used in the bacterial  
CC expression of human Asp2L. The invention relates to a protease (e.g.  
CC Asp2) capable of cleaving the beta secretase site of amyloid precursor  
CC protein (APP). The protease contains a sequence encoding the amino acid  
CC sequence DTG and a sequence encoding DSG or DTG separated by 100-300  
CC amino acids. When mutated the APP gene causes an autosomal dominant form  
CC of Alzheimer's disease. APP localises to the cell surface membrane and  
CC have a single C-terminal transmembrane domain. Proteolytic processing of  
CC APP produces the amyloid beta protein, which is possibly very important  
CC in Alzheimer's disease. The invention includes a nucleotide sequence  
CC encoding the protease, a vector containing the nucleotide sequence, and a  
CC cell line comprising the vector. Methods for screening for inhibitors of  
CC beta secretase activity are also given in the invention. The human  
CC aspartase protein and nucleotide sequences and the methods for  
CC identifying inhibitors of the protease, are useful in the treatment of  
CC Alzheimer's disease  
XX

SQ Sequence 1278 BP; 284 A; 356 C; 353 G; 285 T; 0 U; 0 Other;

Alignment Scores:

Pred. No.: 4.8e-14 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 3 Gaps: 0

US-10-726-967A-52 (1-28) x AAS11714 (1-1278)

QY 1 G|Y|T|T|Y|V|A|G|L|W|E|T|H|T|V|A|G|L|Y|S|E|R|P|R|O|G|I|N|T|H|L|E|U|A|S|N|I|L|E|U|V|A|A|S|P 20  
DB 136 G|G|C|T|A|C|T|A|C|G|T|G|A|G|A|T|G|A|C|C|G|T|G|G|C|A|G|C|C|C|C|G|C|A|G|A|C|G|C|T|C|A|C|T|C|T|G|T|G|A|T 195

QY 21 T|H|G|I|S|E|R|S|E|R|A|S|P|H|E|A|I|A|V|A| 28  
DB 196 A|C|A|G|G|C|A|G|C|A|G|T|A|C|T|T|G|C|A|G|T|G 219

RESULT 2

AAS11714  
ID AAS11714 standard; DNA; 1278 BP.  
XX

AC AAS11714;  
XX

DT 11-SEP-2003 (revised)  
DT 24-OCT-2001 (first entry)  
XX

DE DNA encoding T7-caspase-caspase 8-human aspartyl protease 2a deltaTM.

XX Human; aspartyl protease 1; Asp-1; neutrotropic; neutrotrophic;  
KW aspartyl protease 2; Asp2; amyloid protein precursor; APP;  
KW beta-secretase; Alzheimer's disease; ds.  
XX

OS Homo sapiens.  
OS Enterobacteria phage T7.  
XX

XX Key Location/Qualifiers  
FT CDS 1..1278  
FT /tag= a  
FT /product= "T7-caspase-caspase 8-Aspartyl protease-2a  
FT delta TM"  
XX

MO200149097-A2.

12-JUL-2001.  
XX

PF 09-MAY-2001; 2001WO-IB000797.  
XX  
XX 09-MAY-2001; 2001WO-IB000797.  
XX

PA (BIEN/) BIENKOWSKI M J.  
PA (GURNEY) GURNEY M E.  
PA (HEIN/) HEINRIKSON R L.  
PA (PARO/) PARODI L A.  
PA (YANR/) YAN R.  
XX

PI Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;  
DR WPI; 2001-502548/55.  
DR P-PSDB; AAD07214.  
XX

PT Novel purified polypeptide comprising fragment of mammalian aspartyl  
PT protease 2, lacking Asp2 transmembrane domain and retaining beta  
PT secretase activity of Asp2 useful for identifying inhibitors of Asp2  
PT activity.  
XX

Example 9; Page 158; 185pp; English.

XX The invention relates to a novel purified polypeptide comprising a  
CC fragment of mammalian aspartyl protease 2 (Asp2) protein which lacks the  
CC Asp2 transmembrane domain and the Asp2 protein, and where the polypeptide  
CC and the fragment retain the beta-secretase activity of the mammalian Asp2  
CC protein. Also included is an isoform of amyloid protein precursor (APP)  
CC comprising the amino acid sequence of a APP or its fragment containing an  
CC APP cleavage site recognizable by a mammalian beta-secretase, and further  
CC comprising two lysine residues at the carboxyl terminus of the amino acid  
CC sequence of the mammalian APP or APP fragment. The polypeptides are used  
CC for assaying for modulators of beta-secretase activity; identifying  
CC agents that inhibit the APP processing activity of human Asp2 aspartyl  
CC protease (Hu-Asp2); identifying agents that modulate the activity of Asp2  
CC; and for reducing cellular production of amyloid beta (Abeta) from APP.  
CC Agents identified by the above methods are useful for treating  
CC Alzheimer's disease; and for identifying modulators of amyloid-beta  
CC (Abeta) peptide production, for use in designing therapeutics for the  
CC treatment or prevention of Alzheimer's disease. Probes and primers  
CC derived from Asp nucleic acid sequences are useful for detecting Hu-Asp  
CC nucleic acids in vitro assays and in Northern and Southern blots. The  
CC present sequence represents the coding sequence of T7-caspase-caspase 8-  
CC human-Asp-2a delta TM construct which has a T7 tag, a caspase 8 leader  
CC sequence and cleavage site, and lacks the transmembrane domain. This  
CC construct was used for bacterial expression and purification of human  
CC Asp2a. (Updated on 11-SEP-2003 to standardise OS field)  
XX

SQ Sequence 1278 BP; 284 A; 356 C; 353 G; 285 T; 0 U; 0 Other;

Alignment Scores:

Pred. No.: 4.8e-14 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 4 Gaps: 0

US-10-726-967A-52 (1-28) x AAS11714 (1-1278)

QY 1 G|Y|T|T|Y|V|A|G|L|W|E|T|H|T|V|A|G|L|Y|S|E|R|P|R|O|G|I|N|T|H|L|E|U|A|S|N|I|L|E|U|V|A|A|S|P 20  
DB 136 G|G|C|T|A|C|T|A|C|G|T|G|A|G|A|T|G|A|C|C|G|T|G|G|C|A|G|C|C|C|C|G|C|A|G|A|C|G|C|T|C|A|C|T|C|T|G|T|G|A|T 195

QY 21 T|H|G|I|S|E|R|S|E|R|A|S|P|H|E|A|I|A|V|A| 28  
DB 196 A|C|A|G|G|C|A|G|C|A|G|T|A|C|T|T|G|C|A|G|T|G 219

RESULT 3

AAD17877  
ID AAD17877 standard; cDNA; 1278 BP.  
XX

AC AAD17877;  
XX

DT 10-DEC-2001 (first entry)  
 XX T7-Caspase-Caspase 8 cleavage-human-pro-Asp2(a) lacking TM domain cDNA.  
 DE  
 XX Human; aspartyl protease 1; Aspl1; amyloid precursor protein; APP;  
 XX Alzheimer's disease; AD; dementia; neurofibrillary tangle; gliosis;  
 KW amyloid plaque; neuronal loss; proteolytic; neurotrophic; neuroprotective;  
 KW T7-Caspase-Caspase 8 cleavage-human-pro-Asp 2(a) protein; ss.  
 XX  
 OS Homo sapiens.  
 OS Synthetic.  
 XX  
 XX Key Location/Qualifiers  
 FT CDS 1..1278  
 FT /\*tag= a  
 FT /product= "T7-Caspase-Caspase 8 cleavage-Human-pro- Asp  
 FT 2(a) protein lacking transmembrane domain"  
 XX  
 XX GB2357767-A.  
 XX  
 PD 04-JUL-2001.  
 XX  
 PF 22-SEP-2000; 2000GB-00023315.  
 XX  
 PR 23-SEP-1999; 99US-00404133.  
 PR 23-SEP-1999; 99US-0155493P.  
 PR 23-SEP-1999; 99WO-US020881.  
 PR 13-OCT-1999; 99US-00416901.  
 PR 06-DEC-1999; 99US-0169232P.  
 XX  
 PA (PHMA ) PHARMACIA & UPJOHN CO.  
 XX  
 PI Bienkowski MJ, Gurney M;  
 XX  
 DR WPI: 2001-444208/48.  
 DR P-PSDB; AAE10641.  
 XX  
 PT Polypeptide comprising fragments of human aspartyl protease with amyloid  
 PT precursor protein processing activity and alpha-secretase activity, for  
 PT identifying modulators useful in treating Alzheimer's disease.  
 XX  
 PS Example 9; Page 128; 187pp; English.  
 XX  
 CC The patent discloses human aspartyl protease 1 (hu-Asp1) or modified Asp1  
 CC proteins which lack transmembrane domain or amino terminal domain or  
 CC cytoplasmic domain and retains alpha-secretase activity and amyloid  
 CC protein precursor (APP) processing activity. The proteins of the  
 CC invention are useful for assaying hu-Asp1 alpha-secretase activity, which  
 CC in turn is useful for identifying modulators of hu-Asp1 alpha-secretase  
 CC activity, where modulators that increase hu-Asp1 alpha-secretase activity  
 CC are useful for treating Alzheimer's disease (AD) which causes progressive  
 CC dementia with consequent formation of amyloid plaques, neurofibrillary  
 CC tangles, gliosis and neuronal loss. Hu-Asp1 protease substrate is useful  
 CC for assaying hu-Asp1 proteolytic activity, by contacting hu-Asp1 protein  
 CC with the substrate under acidic conditions and determining the level of  
 CC hu-Asp1 proteolytic activity. The present sequence is a cDNA encoding T7-  
 CC Caspase-Caspase 8 cleavage-human-pro-Asp 2(a) protein lacking a  
 CC transmembrane (TM) domain which is generated from human Asp 2(a) protein  
 CC by the addition of T7 tag and caspase 8 leader sequence at its N-terminal  
 CC end and deletion of its C-terminal transmembrane domain  
 XX  
 SQ Sequence 1278 BP; 284 A, 356 C, 353 G, 285 T; 0 U; 0 Other;  
 XX  
 Alignment Scores:  
 Pred. No.: 4,8e-14 Length: 1278  
 Score: 144.00 Matches: 28  
 Percent Similarity: 100.00% Conservative: 0  
 Best Local Similarity: 100.00% Mismatches: 0  
 Query Match: 100.00% Indels: 0  
 DB: 4 Gaps: 0  
 US-10-726-967A-52 (1-28) x AAD17877 (1-1278)

QY 1 GYTYTYTYValGlumetThrValGlySerProProGlnThrLeuAsnIleLeuValAsp 20  
 |||||  
 DB 136 GGCTACTGAGTGGAGATGACGCTGGCGACGCCCGCAGACGCTCAACATCCTGGTGAT 195  
 |||||  
 QY 21 ThrGlySerSerAspPheAlaVal 28  
 |||||  
 DB 196 ACAGCGACGACGACTGACTTGCACGTG 219  
 |||||  
 RESULT 4  
 AAD13033  
 ID AAD13033 standard; cDNA; 1278 BP.  
 XX  
 XX AAD13033;  
 XX  
 XX 23-OCT-2001 (first entry)  
 XX  
 DE T7-Caspase-Caspase 8 cleavage-Human-pro-Asp2(a) deltatm protein cDNA.  
 XX  
 KW Human; aspartyl protease 2a; Asp 2a; beta-amyloid precursor protein; APP;  
 KW beta-secretase; Alzheimer's disease; dementia; amyloid plaque; gliosis;  
 KW neurofibrillary tangle; neuronal loss; amyloid-beta peptide; neurotrophic;  
 KW neuroprotective; antisense therapy; gene therapy;  
 KW caspase-caspase 8 cleavage-pro-Asp2(a) deltatm protein; ss.  
 XX  
 OS Homo sapiens.  
 OS Synthetic.  
 XX  
 XX Key Location/Qualifiers  
 FH CDS 1..1278  
 FT /\*tag= a  
 FT /product= "T7-Caspase-Caspase 8 cleavage-Human-pro-  
 FT Asp2(a) deltatm protein"  
 XX  
 XX W0200150829-A2.  
 XX  
 PD 19-JUL-2001.  
 XX  
 PF 09-MAY-2001; 2001WO-1B000799.  
 XX  
 PR 09-MAY-2001; 2001WO-1B000799.  
 XX  
 PA (BIEN/) BIENKOWSKI M J.  
 PA (GURN/) GURNEY M B.  
 PA (HEIN/) HEINRIKSON R L.  
 PA (PARO/) PARODI L A.  
 PA (YANR/) YAN R.  
 XX  
 PI Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;  
 XX  
 DR WPI: 2001-483072/52.  
 DR P-PSDB; AAE06871.  
 XX  
 PT Novel purified polypeptide comprising fragment of mammalian aspartyl  
 PT protease 2, lacking Asp2 transmembrane domain and retaining beta  
 PT secretase activity of Asp2 useful for identifying inhibitors of Asp2  
 PT activity.  
 XX  
 PS Example 9; Page 158; 185pp; English.  
 XX  
 CC The invention relates to human aspartyl proteases (Hu-Asp), beta-amyloid  
 CC precursor protein (APP) isoforms and their corresponding DNA molecules.  
 CC Human aspartyl proteases can act as beta-secretase proteases useful for  
 CC treating Alzheimer's disease. APP isoforms are useful for identifying  
 CC modulators of amyloid-beta peptide production, for use in designing  
 CC therapeutics for the treatment and prevention of Alzheimer's disease,  
 CC dementia, formation of amyloid plaques, neurofibrillary tangles, gliosis  
 CC and neuronal loss. APP isoforms are also used in methods for identifying  
 CC inhibitors and modulators of human Asp2 activity. The invention relates  
 CC to a method for identifying agents that modulate the activity of human  
 CC aspartyl protease Asp2. Amyloid-beta peptides obtained from APP are used  
 CC as a means to screen in cellular assays for the inhibitors of beta- and  
 CC gamma- secretase. Hu-Asp DNA fragments are useful as probes or primers in  
 CC polymerase chain reactions (PCR). The probes are useful for detecting Hu-

CC Asp nucleic acids in in vitro assays and in Northern and Southern blots.  
CC The present cDNA sequence encodes T7-Caspase-Caspase 8 cleavage- Human-  
CC pro-aspartyl protease 2a (Asp2a) delta TM protein which is obtained by the  
CC addition of T7 tag and caspase 8 leader sequence at the N-terminal end  
CC and deletion of the transmembrane domain at the C-terminal end of Hu-  
CC Asp2a. Human Asp2a has beta-secretase activity

SQ Sequence 1278 BP; 284 A; 356 C; 353 G; 285 T; 0 U; 0 Other;

Alignment Scores:

Pred. No.:	4.8e-14	Length:	1278
Score:	144.00	Matches:	28
Percent Similarity:	100.00%	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	100.00%	Indels:	0
DB:	4	Gaps:	0

US-10-726-967a-52 (1-28) x AAD06751 (1-1278)

QY 1 G|Y|Y|T|Y|V|A|G|U|M|E|T|T|H|V|A|G|I|S|E|P|P|R|O|G|I|N|T|H|L|E|U|E|N|I|L|E|U|V|A|A|P 20

Db 136 GGCCTACTACCTGGAGATGACCGTGGCGAGCCCGCCGAGACGCTCAACATCCTGTGGAT 195

QY 21 ThrGlySerSerAspAspAspVal 28

Db 196 ACAAGCAGCAGTAACCTTTCAGTG 219

RESULT 5

ID AAD06751 standard; cDNA; 1278 BP.

AC AAD06751;

DT 10-AUG-2001 (first entry)

DE T7-Caspase-Caspase 8-cleavage-human-pro-Asp-2(a) delta TM protein CDNA.

KM Human; alpha-secretase; amyloid precursor protein; APP; therapy;

KW Alzheimer's disease; anti-Alzheimer's; aspartyl protease 2a; Asp2a;

OS Homo sapiens.

OS Synthetic.

Key Location/Qualifiers

CDS 1..1278  
/tag= a  
/product= "T7-Caspase-Caspase 8-cleavage-human-pro- Asp-  
2(a) delta TM protein"

XX WO200123533-A2.

PD 05-APR-2001.

PF 22-SEP-2000; 2000WO-US026080.

PR 23-SEP-1999; 99US-0155493P.

PR 13-SEP-1999; 99WO-US020881.

PR 13-OCT-1999; 99US-00416901.

PR 06-DEC-1999; 99US-0169232P.

PA (PHAA ) PHARMACIA & UPJOHN CO.

PI Gurney M, Bienkowski MJ;

XX WPI; 2001-290516/30.

XX P-PSDB; AAB02593.

PT Enzymes that cleave the alpha-secretase site of the amyloid precursor  
XX protein, useful for the treatment of Alzheimer's disease.  
XX Example 9; Page 157; 189pp; English.

CC The present invention relates to enzymes for cleaving the alpha-  
CC secretase site of the amyloid precursor protein (APP) and methods of  
CC identifying those enzymes. The methods may be used to identify enzymes  
CC that may be used to cleave the alpha-secretase cleavage site of the APP  
CC protein. The enzymes may be used to treat or modulate the progress of  
CC Alzheimer's disease. The present sequence is a cDNA encoding human  
CC aspartyl protease 2a (Asp-2a) caspase-caspase 8-delta TM protein which is  
CC obtained by deleting the transmembrane domain and adding a T7-caspase  
CC leader sequence at the N-terminal end. This sequence has beta-secretase  
XX protease activity

SQ Sequence 1278 BP; 284 A; 356 C; 353 G; 285 T; 0 U; 0 Other;

Alignment Scores:	
Pred. No.:	4.8e-14
Score:	144.00
Percent Similarity:	100.00%
Best Local Similarity:	100.00%
Query Match:	100.00%
DB:	4
Length:	1278
Matches:	28
Conservative:	0
Mismatches:	0
Indels:	0
Gaps:	0

US-10-726-967a-52 (1-28) x AAD06751 (1-1278)

QY 1 G|Y|Y|T|Y|V|A|G|U|M|E|T|T|H|V|A|G|I|S|E|P|P|R|O|G|I|N|T|H|L|E|U|E|N|I|L|E|U|V|A|A|P 20

Db 136 GGCCTACTACCTGGAGATGACCGTGGCGAGCCCGCCGAGACGCTCAACATCCTGTGGAT 195

QY 21 ThrGlySerSerAspAspAspVal 28

Db 196 ACAAGCAGCAGTAACCTTTCAGTG 219

RESULT 6

ID AAS11529 standard; cDNA; 1278 BP.

AC AAS11529;

DT 24-OCT-2001 (first entry)

DE T7-Caspase-caspase-8-Human-pro-Asp 2(a) delta TM fusion protein CDNA.

KM Human; Aspartyl protease; beta-secretase; neurotrophic; ASP2;

KW neuroprotective; amyloid protein precursor; APP; Alzheimer's disease;

KW amyloid-beta; Abeta; T7-Caspase-caspase 8-Human-pro-Asp 2(a) delta TM;

OS se.

OS Homo sapiens.

OS Synthetic.

Key Location/Qualifiers

CDS 1..1278  
/tag= a  
/product= "T7-Caspase-caspase 8-Human-pro-Asp 2(a) delta  
TM fusion protein"

FT sig\_peptide 43..87

FT mat\_peptide 88..1275

FT /note= "Caspase leader sequence"

FT /label= Mature Asp 2(a)

FT /note= "Also encodes 5 extra N-terminal amino acids  
constituting a caspase 8 cleavage site"

XX WO200149098-A2.

PD 12-JUL-2001.

PF 09-MAY-2001; 2001WO-IB000798.

PR 09-MAY-2001; 2001WO-IB000798.

PA (BIEN/) BIENKOWSKI M J.

PA (GURN/) GURNEY M E.

PA (HEIN/) HEINRIKSON R L.

PA (PARO/) PARODI L A.  
 PA (YANR/) YAN R.  
 XX  
 XX  
 PI Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;  
 XX  
 XX WPI: 2001-502549/55.  
 DR P-PSDB; AAU06615.  
 XX  
 XX Novel purified polypeptide comprising fragment of mammalian aspartyl  
 PT protease 2, lacking Asp2 transmembrane domain and retaining beta  
 PT secretase activity of Asp2 useful for identifying inhibitors of Asp2  
 PT activity.  
 XX  
 XX Example 9, Page 158, 185pp; English.  
 PS  
 XX  
 XX The invention relates to a purified polypeptide comprising a fragment of  
 CC mammalian aspartyl protease (Asp) 2 protein which lacks the Asp2  
 CC transmembrane domain and the Asp2 protein, and where the polypeptide and  
 CC the fragment retain the beta-secretase activity of the mammalian Asp2  
 CC protein. The invention also details polynucleotides for the Asp proteins  
 CC and vectors expressing them, and a polypeptide (isoform of amyloid  
 CC protein precursor (APP)) comprising the amino acid sequence of an APP or  
 CC its fragment containing an APP cleavage site recognizable by a mammalian  
 CC beta-secretase, and further comprising two lysine residues at the  
 CC carboxyl terminus of the amino acid sequence of the mammalian APP or APP  
 CC fragment. Also included in the invention are methods of identifying  
 CC modulators or inhibitors of Asp2. Modulators and inhibitors of Asp2 are  
 CC useful for treating Alzheimer's disease. APP is useful in methods for  
 CC identifying inhibitors or modulators of human Asp2 activity and amyloid-  
 CC beta (Abeta) peptide production. APP is also useful in designing  
 CC therapeutics for the treatment or prevention of Alzheimer's disease. APP  
 CC comprising the APP-Sw-beta-secretase peptide sequence (NLBD), which is  
 CC associated with increased levels of Abeta processing is useful in assays  
 CC relating the Alzheimer's research. The expression vector is useful for  
 CC recombinantly expressing APP. Nucleic acids that hybridize to Asp  
 CC oligonucleotides are useful as probes or primers. The probes are useful  
 CC for detecting hu-Asp nucleic acids in in vitro assays and in Northern and  
 CC Southern blots. The present sequence encodes T7-caspase-8 human-  
 CC pro-Asp 2(a) delta TM fusion protein which has a N-terminal T7 tag to aid  
 CC purification when expressed in E. coli, a Caspase leader sequence and a  
 CC caspase 8 cleavage signal to aid cleavage of the signal peptide  
 CC  
 SQ Sequence 1278 BP; 284 A; 356 C; 353 G; 285 T; 0 U; 0 Other;  
 XX  
 XX Alignment Scores:  
 Pred. No.: 4.8e-14 Length: 1278  
 Score: 144.00 Matches: 28  
 Percent Similarity: 100.00% Conservative: 0  
 Best Local Similarity: 100.00% Mismatches: 0  
 Query Match: 100.00% Indels: 0  
 DB: 4 Gaps: 0  
 US-10-726-967A-52 (1-28) x ABL52469 (1-1278)  
 QY 1 G|Y|T|Y|T|Y|A|G|L|u|e|c|T|h|V|A|G|I|S|e|P|P|o|G|I|n|T|h|r|e|u|s|e|n|l|e|u|V|a|A|s|p 20  
 DB 136 G|G|C|T|A|C|T|G|A|G|A|T|G|A|C|C|G|G|G|G|C|A|G|C|C|C|C|G|C|A|G|A|C|G|T|C|A|C|A|T|C|T|G|T|G|A|T 195  
 QY 21 T|h|G|I|S|e|S|e|r|S|e|r|A|s|p|h|e|a|l|A|a|l 28  
 DB 196 A|C|A|G|G|C|A|G|C|A|G|A|C|T|T|G|C|A|G|T|G 219  
 XX  
 XX RESULT 7  
 ABL52469  
 ID ABL52469 standard; cDNA; 1278 BP.  
 AC ABL52469;  
 XX  
 XX 16-JUL-2002 (first entry)  
 DT  
 XX  
 XX T7-caspase-caspase 8 cleavage-human-pro-Asp-2(a)deltaTM nucleotide.  
 DE  
 XX Human; Asp-1; Asp-2; aspartyl protease; Alzheimer's disease; proteolytic;

KM amyloid precursor protein; APP; gene; ss.  
 XX  
 XX Homo sapiens.  
 OS  
 XX  
 XX Key Location/Qualifiers  
 FH CDS 1..1278  
 FT 1..1278  
 FT /\*tag= a  
 FT /product= "T7-caspase-caspase 8 cleavage-human-pro- Asp-  
 FT 2(a)deltaTM"  
 XX  
 XX GB2367060-A.  
 PN  
 XX  
 XX 27-MAR-2002.  
 PD  
 XX  
 XX 29-OCT-2001; 2001GB-00025934.  
 PF  
 XX  
 XX 23-SEP-1999; 99US-00404133.  
 PR 23-SEP-1999; 99US-0155493P.  
 PR 23-SEP-1999; 99WO-US020881.  
 PR 13-OCT-1999; 99US-00416901.  
 PR 06-DEC-1999; 99US-0169232P.  
 PR 22-SEP-2000; 2000GB-00023315.  
 XX  
 XX (PHAA ) PHARMACIA & UPJOHN CO.  
 PA  
 XX  
 XX Bienkowski MJ, Gurney M;  
 PI  
 XX  
 XX WPI: 2002-397167/43.  
 DR P-PSDB; ABB78602.  
 DR  
 XX  
 XX Human aspartyl protease 1 substrates useful in assays to detect aspartyl  
 PT protease activity, e.g. for the diagnosis of Alzheimer's disease.  
 PT  
 XX  
 XX Example 9, Page 128, 182pp; English.  
 PS  
 XX  
 XX The present invention describes a human aspartyl protease 1 (hu-Asp1)  
 CC substrate (I) which comprises a peptide of no more than 50 amino acids,  
 CC and which comprises the 8 amino acid sequence Gly-Leu-Ala-Leu-Ala-Leu-  
 CC Glu-Pro. Also described are: (1) a method (II) for assaying hu-Asp1  
 CC proteolytic activity, comprising: (a) contacting a hu-Asp1 protein with  
 CC (1) under acidic conditions; and (b) determining the level of hu-Asp1  
 CC proteolytic activity; (2) a purified polynucleotide (III) comprising a  
 CC nucleotide sequence that hybridizes under stringent conditions to the non-  
 CC coding strand complementary to a defined 1804 nucleotide sequence (see  
 CC ABL52456) where the nucleotide sequence encodes a polypeptide having Asp1  
 CC proteolytic activity and lacks nucleotides encoding a transmembrane  
 CC domain; (3) a purified polynucleotide (III') comprising a sequence that  
 CC hybridizes under stringent conditions to (III) (the nucleotide sequence  
 CC encodes a polypeptide further lacking a pro-peptide domain corresponding  
 CC to amino acids 23-62 of hu-Asp1 (see ABB78589); (4) a vector (IV)  
 CC comprising (III) or (III'); and (5) a host cell (V) transformed or  
 CC transfected with (III), (III') and/or (IV). The hu-Asp1 protease  
 CC substrate (I) may be used as an enzyme substrate in assays to detect  
 CC aspartyl protease activity, (II) and therefore diagnose diseases  
 CC associated with aberrant hu-Asp1 expression and activity such as  
 CC Alzheimer's disease. Hu-Asp1 has been localised to chromosome 21, while  
 CC hu-Asp2 has been localised to chromosome 11q23.3-24.1. The present  
 CC sequence encodes T7-caspase-caspase 8 cleavage-human-pro-Asp-2(a)deltaTM,  
 CC which is given in an example from the present invention  
 CC  
 XX  
 SQ Sequence 1278 BP; 284 A; 356 C; 353 G; 285 T; 0 U; 0 Other;  
 XX  
 XX Alignment Scores:  
 Pred. No.: 4.8e-14 Length: 1278  
 Score: 144.00 Matches: 28  
 Percent Similarity: 100.00% Conservative: 0  
 Best Local Similarity: 100.00% Mismatches: 0  
 Query Match: 100.00% Indels: 0  
 DB: 6 Gaps: 0  
 US-10-726-967A-52 (1-28) x ABL52469 (1-1278)  
 QY 1 G|Y|T|Y|T|Y|A|G|L|u|e|c|T|h|V|A|G|I|S|e|P|P|o|G|I|n|T|h|r|e|u|s|e|n|l|e|u|V|a|A|s|p 20

```

Db      136  |||GGCTACTACGTGAGATGACCGTGGCGACCCCGCGACGCTCAACATCTGTGAT 195
Qy      21  |||ThrglySerSerAsenPheAlaVal 28
Db      196  |||ACAGGACAGCACTTGTGCAGTG 219

RESULT 8
ADJ94339 ADJ94339 standard, cDNA, 1278 BP.
AC      ADJ94339;
DT      03-JUN-2004 (first entry)
DE      Human T7-Caspase-Caspase 8 cleavage-human-pro-Asp-2(a)deltaTM cDNA.
XX
XX      Human; sg, gene; aspartyl protease; Asp-1; Asp-2(a); Asp-2(b);
KW      beta secretase; amyloid protein precursor; APP; Alzheimer's disease;
KW      neurotrophic; neuroprotective; amyloid beta; mutant.
OS      Homo sapiens.
OS      Synthetic.
XX
PN      US6706485-B1.
XX
PD      16-MAR-2004.
XX
PF      12-APR-2000; 2000US-00548376.
XX
PR      24-SEP-1998; 98US-0101594P.
PR      23-SEP-1999; 99US-00404133.
PR      23-SEP-1999; 99US-0155493P.
PR      23-SEP-1999; 99WO-US020881.
PR      13-OCT-1999; 99US-00416901.
XX
PA      (PHAA ) PHARMACIA & UPJOHN CO.
XX
XX      Gurney ME, Blenkowski MJ, Heinrikson RL, Parodi LA, Yan R;
XX      WPI; 2004-236722/22.
XX      P-PSDB; ADJ94340.
XX
PT      Identifying agents that modulate activity of Asp2 aspartyl protease
PT      useful for treating or preventing Alzheimer's disease involves comparing
PT      APP processing activity of protease in presence and absence of test
PT      agent.
XX
PS      Disclosure; SEQ ID NO 27; 109pp; English.
XX
XX      The invention relates to identifying agents that modulate activity of
XX      Asp2 (e.g. a beta-secretase, e.g. human Asp-2(b) appearing as ID 6,
XX      encoded by ID 5) aspartyl protease, involves contacting Asp2 with amyloid
XX      precursor protein (APP) in the presence and absence of a test agent,
XX      where Asp2 is a recombinant polypeptide and processes APP into amyloid
XX      beta, determining APP processing activity of Asp2 in presence and absence
XX      of the test agent, and comparing the activities to identify agents that
XX      modulate the activity of Asp2. Also disclosed are the cDNA and proteins
XX      for human Asp-1 and Asp-2(a), mouse Asp-2(b), a vector comprising the
XX      nucleic acid encoding Hu-Asp2 protease sequence, a host cell comprising
XX      the vector and the method of producing Hu-Asp polypeptide, an isolated
XX      antibody that specifically binds to Hu-Asp polypeptides, identifying a
XX      cell that can be used to screen for inhibitors of beta secretase
XX      activity, novel isoforms of amyloid protein precursor (APP), where the
XX      last 2 carboxy terminus amino acids of that isoform are both lysine
XX      mutation where KM at 595-596 is mutated to NV, designated e.g. APP695-SW
XX      or APP695-SW-KK, or a V to F mutation at 642, e.g. APP695-VF, all useful
XX      for assaying for beta secretase activity and screening for inhibitors of
XX      beta-secretase) and polynucleotides that encode the APP proteins. The
XX      method is useful for identifying agents that modulate the activity
XX      (amyloid precursor protein processing activity) of Asp2 aspartyl
XX      protease. Preferably, the method is useful for identifying agents that

```

```

CC      inhibit Asp2 aspartyl protease activity. The inhibitors of amyloid
CC      precursor protein processing, are useful for treating or preventing
CC      Alzheimer's disease. The present sequence encodes an aspartyl protease
CC      mutant construct (e.g. lacking a transmembrane domain and/or including a
CC      caspase cleavage site) used to investigate the cleavage activity of Asp2
CC      proteins.
XX
SQ      Sequence 1278 BP; 284 A; 356 C; 353 G; 285 T; 0 U; 0 Other;

Alignment Scores:
Pred. No.:      4.8e-14
Score:          144.00
Percent Similarity: 100.00%
Best Local Similarity: 100.00%
Query Match:    100.00%
DB:             12
Gaps:           0

US-10-726-967A-52 (1-28) x ADJ94339 (1-1278)
Qy      1  |||GlyTyrTyrValGluMetThrValGlySerProProGlnThrLeuAniLeuValAsp 20
Db      136  |||GGCTACTACGTGAGATGACCGTGGCGACCCCGCGACGCTCAACATCTGTGAT 195
Qy      21  |||ThrglySerSerAsenPheAlaVal 28
Db      196  |||ACAGGACAGCACTTGTGCAGTG 219

RESULT 9
ADJ94339 ADJ94339 standard, cDNA, 1278 BP.
ID      ADJ94339;
XX
AC      ADJ94339;
XX
XX      29-JUN-2004 (first entry)
XX
DE      T7-Caspase-Caspase 8 cleavage-Human-pro-Asp-2(a)deltaTM DNA.
XX
XX      Aspartyl protease; Asp; beta secretase; amyloid precursor protein; APP;
KW      Alzheimer's disease; gene therapy; caspase; human; gene; chimeric; ds.
XX
XX      Homo sapiens.
OS      Chimeric.
OS      Unidentified.
XX
XX      Key      Location/Qualifiers
XX      CDS      1..1278
XX              /tag=a
XX              /product="T7-Caspase-Caspase 8 cleavage-Human-pro-Asp-
XX              2(a)deltaTM protein"
XX
XX      US6737510-B1.
XX
XX      18-MAY-2004.
XX
XX      12-APR-2000; 2000US-00548373.
XX
XX      24-SEP-1998; 98US-0101594P.
XX      23-SEP-1999; 99US-00404133.
XX      23-SEP-1999; 99US-0155493P.
XX      23-SEP-1999; 99WO-US020881.
XX      13-OCT-1999; 99US-00416901.
XX
XX      (PHAA ) PHARMACIA & UPJOHN CO.
XX
XX      Gurney ME, Blenkowski MJ, Heinrikson RL, Parodi LA, Yan R;
XX      WPI; 2004-387112/36.
XX      P-PSDB; ADJ94336.
XX
XX      New Asp2 aspartyl protease protein comprising tripeptides DTG and DSG
XX      involved in processing amyloid precursor protein into amyloid beta,
XX      useful in preparing a composition for treating or preventing Alzheimer's
XX      disease.

```

XX Example 9; SEQ ID NO 27; 108bp; English.

XX The invention relates to a method for identifying an agent that decreases

XX the protease activity of the aspartyl protease (Asp) polypeptide. It also

XX provides enzyme and enzymatic procedures for cleaving the beta secretase

XX cleavage site of the amyloid precursor protein (APP). The invention is

XX useful in preparing a composition for treating or preventing Alzheimer's

XX disease. It is also useful in gene therapy. The present sequence is T7-

XX Caspase-Caspase 8 cleavage-Human-pro-Asp-2(a)deltaTM chimeric DNA. This

XX sequence is used to illustrate the method of the invention.

XX

XX Sequence 1278 BP; 284 A; 356 C; 353 G; 285 T; 0 U; 0 Other;

XX

XX Alignment Scores:

XX Pred. No.: 4,8e-14 Length: 1278

XX Score: 144.00 Matches: 28

XX Percent Similarity: 100.00% Conservativeness: 0

XX Best Local Similarity: 100.00% Mismatches: 0

XX Query Match: 100.00% Indels: 0

XX DB: 12 Gaps: 0

XX

XX US-10-726-967a-52 (1-28) x ADO50435 (1-1278)

XX

XX 1 GYTYTYTYValGluMetThrValGlySerProProGlnThrLeuAniLeuValAsp 20

XX 136 GGCTACTACGTGAGATGACCGTGGGACGCCGCCGACGCTCAACATCTGTGAT 195

XX

XX 21 ThrGlySerSerAspAhpheAlaVal 28

XX 196 ACAGGACGACGTAACCTTTCACAGT 219

XX

XX RESULT 10

XX ADR75348

XX ID ADR75348 standard; DNA; 1278 BP.

XX

XX ADR75348;

XX

XX 18-NOV-2004 (first entry)

XX

XX T7-Caspase-Caspase 8 cleavage-Human-pro-Asp-2(a)deltaTM DNA.

XX

XX Aspartyl protease; Asp; amyloid precursor protein; APP; amyloid beta;

XX chromosome identification; Alzheimer's disease; human; caspase; chimeric;

XX gene; de.

XX

XX Homo sapiens.

XX

XX Chimeric.

XX

XX Unidentified.

XX

XX Key Location/Qualifiers

XX FT CDS 1..1278

XX FT /product= "T7-Caspase-Caspase 8 cleavage-Human-pro-Asp-

XX FT 2(a)deltaTM protein"

XX

XX US2004166507-A1.

XX

XX 26-AUG-2004.

XX

XX 29-AUG-2003; 2003US-00652045.

XX

XX 24-SEP-1998; 98US-0101594P.

XX

XX 23-SEP-1999; 99US-00404133.

XX

XX 13-SEP-1999; 99US-0155493P.

XX

XX 13-OCT-1999; 99US-00416901.

XX

XX (GURN/) GURNEY M. E.

XX

XX (BIEN/) BIENKOWAKI M. J.

XX

XX (HEIN/) HEINRIKSON R. L.

XX

XX (PARO/) PARODI L. A.

XX

XX (YANR/) YAN R.

PI Gurney ME, Bienkowiak MJ, Heinrichson RL, Parodi LA, Yan R;

XX MPI; 2004-624916/60.

XX

XX P-PDB; ADR75349.

XX

XX Novel purified/isolated polynucleotide encoding polypeptide having

XX aspartyl protease activity involved in processing amyloid precursor

XX protein into amyloid beta, useful in identifying agent decreasing

XX activity of aspartyl protease.

XX

XX Example 9; SEQ ID NO 27; 107bp; English.

XX

XX The invention relates to nucleic acid sequences encoding aspartyl

XX protease (Asp) polypeptides having aspartyl protease activity involved in

XX processing amyloid precursor protein (APP) into amyloid beta. The

XX invention also relates to a method for identifying an agent that

XX decreases the protease activity of the Asp. Asp DNA is useful in

XX chromosome identification as they can hybridise with a specific location

XX on a human chromosome and in identifying the relationship between genes

XX and diseases (particular gene responsible for causing diseases). It is

XX also useful for identifying candidates to modulate the progression of

XX Alzheimer's disease. Asp is useful in raising antibodies that are useful

XX in diagnostic assay for detecting Hu-Asp polypeptide expression. The

XX present sequence is the T7-Caspase-Caspase 8 cleavage-Human-pro-Asp-

XX 2(a)deltaTM DNA. This sequence is used to illustrate the method of the

XX invention.

XX

XX Sequence 1278 BP; 284 A; 356 C; 353 G; 285 T; 0 U; 0 Other;

XX

XX Alignment Scores:

XX Pred. No.: 4,8e-14 Length: 1278

XX Score: 144.00 Matches: 28

XX Percent Similarity: 100.00% Conservativeness: 0

XX Best Local Similarity: 100.00% Mismatches: 0

XX Query Match: 100.00% Indels: 0

XX DB: 13 Gaps: 0

XX

XX US-10-726-967a-52 (1-28) x ADR75348 (1-1278)

XX

XX 1 GYTYTYTYValGluMetThrValGlySerProProGlnThrLeuAniLeuValAsp 20

XX 136 GGCTACTACGTGAGATGACCGTGGGACGCCGCCGACGCTCAACATCTGTGAT 195

XX

XX 21 ThrGlySerSerAspAhpheAlaVal 28

XX 196 ACAGGACGACGTAACCTTTCACAGT 219

XX

XX RESULT 11

XX AAD17895

XX ID AAD17895 standard; cDNA; 1287 BP.

XX

XX AAD17895;

XX

XX 10-DEC-2001 (first entry)

XX

XX Human-Asp 2(b) protein lacking transmembrane domain encoding cDNA.

XX

XX Human; aspartyl protease 2b; Asp2b; amyloid precursor protein; APP;

XX Alzheimer's disease; AD; dementia; neurofibrillary tangle; gliosis;

XX amyloid plaque; neuronal loss; proteolytic; neurotrophic; neuroprotective;

XX se.

XX

XX Homo sapiens.

XX

XX Synthetic.

XX

XX Key Location/Qualifiers

XX FT CDS 1..1287

XX FT /product= "Human-Asp 2(b) protein lacking transmembrane

XX FT domain"

XX

XX GB2357767-A.



```

PD 04-JUL-2001.
XX
PF 22-SEP-2000; 2000GB-00023315.
XX
PR 23-SEP-1999; 99US-00404133.
PR 23-SEP-1999; 99US-0155493P.
PR 23-SEP-1999; 99MO-US020881.
PR 13-OCT-1999; 99US-00416901.
PR 06-DEC-1999; 99US-0169232P.
XX
PA (PHAA ) PHARMACIA & UPJOHN CO.
PI Bienkowskei MJ, Gurney M;
DR WPI; 2001-444208/48.
P-PSDB; AAE10646.
XX
PT Polypeptide comprising fragments of human aspartyl protease with amyloid
PT precursor protein processing activity and alpha-secretase activity, for
PT identifying modulators useful in treating Alzheimer's disease.
XX
PS Example 10; Page 137; 187pp; English.
XX
CC The patent discloses human aspartyl protease 1 (hu-Asp1) or modified Asp1
CC proteins which lack transmembrane domain or amino terminal domain or
CC cytoplasmic domain and retains alpha-secretase activity and amyloid
CC protein precursor (APP) processing activity. The proteins of the
CC invention are useful for assaying hu-Asp1 alpha-secretase activity, which
CC in turn is useful for identifying modulators of hu-Asp1 alpha-secretase
CC activity, where modulators that increase hu-Asp1 alpha-secretase
CC are useful for treating Alzheimer's disease (AD) which causes progressive
CC dementia with consequent formation of amyloid plaques, neurofibrillary
CC tangles, gliosis and neuronal loss. Hu-Asp1 protease substrate is useful
CC for assaying hu-Asp1 proteolytic activity, by contacting hu-Asp1 protein
CC with the substrate under acidic conditions and determining the level of
CC hu-Asp1 proteolytic activity. The present sequence is a cDNA encoding
CC human Asp 2(b) protein lacking a transmembrane (TM) domain which is
CC generated by the deletion of the C-terminal TM domain and intracellular
CC domains of human Asp 2(b) protein
XX
SQ Sequence 1287 BP; 271 A; 370 C; 384 G; 262 T; 0 U; 0 Other;
XX
Alignment Scores:
Pred. No.: 4,84e-14 Length: 1287
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
Gaps: 0
US-10-726-967A-52 (1-28) x AAD17895 (1-1287)
XX
QY 1 GlyTYYTYRVALGIuMeTherValGISeSrProProGIInThrlEuAnIlleuValAasp 20
Db 220 GGCTACTACGTGAGATGACCGTGGGACACCCCGCAGACACCTCAACATCTCGTGAGT 279
QY 21 ThrgISeSrSeSrAspnhelAval 28
Db 280 ACAGGACAGCAGTAACCTTTCAGTG 303
RESULT 12
AAD13276
ID AAD13276 standard; cDNA; 1287 BP.
XX
XX AAD13276;
XX AC
XX 23-OCT-2001 (first entry)
XX
XX Human-Asp2(b) deltaTM protein cDNA.
XX
XX Human; aspartyl protease 2b; Asp 2b; beta-amyloid precursor protein; APP;
XX beta-secretase; Alzheimer's disease; dementia; amyloid plaque; gliosis;
XX neurofibrillary tangle; neuronal loss; amyloid-beta peptide; nootropic;

```

KW neuroprotective; antisense therapy; Asp2(b) deltaTM protein;  
 XX gene therapy; ss.  
 XX Homo sapiens.  
 OS Synthetic.  
 XX  
 XX  
 FH Key Location/Qualifiers  
 FT CDS 1..1287  
 FT /\*tag= a  
 FT /product= "Human Asp2(b) deltaTM protein"  
 EN  
 PN MO200150829-A2.  
 PD 19-JUL-2001.  
 XX  
 XX  
 PF 09-MAY-2001; 2001MO-IB000799.  
 PR 09-MAY-2001; 2001MO-IB000799.  
 XX  
 PA (BIEN/) BIENKOWSKI M J.  
 PA (GURN/) GURNEY M E.  
 PA (HEIN/) HEINRIKSON R L.  
 PA (PARO/) PARODI L A.  
 PA (YANR/) YAN R.  
 XX  
 PI Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;  
 DR WPI; 2001-483072/52.  
 DR P-PSDB; AA06891.  
 XX  
 PT Novel purified polypeptide comprising fragment of mammalian aspartyl  
 PT protease 2, lacking Asp2 transmembrane domain and retaining beta  
 PT secretase activity of Asp2 useful for identifying inhibitors of Asp2  
 PT activity.  
 XX  
 PS Example 10; Page 166-167; 185pp; English.  
 XX  
 XX The invention relates to human aspartyl proteases (Hu-Asp), beta-amyloid  
 CC precursor protein (APP) isoforms and their corresponding DNA molecules.  
 CC Human aspartyl proteases can act as beta-secretase proteases useful for  
 CC treating Alzheimer's disease. APP isoforms are useful for identifying  
 CC modulators of amyloid-beta peptide production, for use in designing  
 CC therapeutics for the treatment and prevention of Alzheimer's disease,  
 CC dementia, formation of amyloid plaques, neurofibrillary tangles, gliosis  
 CC and neuronal loss. APP isoforms are also used in methods for identifying  
 CC inhibitors and modulators of human Asp2 activity. The invention relates  
 CC to a method for identifying agents that modulate the activity of human  
 CC aspartyl protease Asp2. Amyloid-beta peptides obtained from APP are used  
 CC as a means to screen in cellular assays for the inhibitors of beta- and  
 CC gamma- secretase. Hu-Asp DNA fragments are useful as probes or primers in  
 CC polymerase chain reactions (PCR). The probes are useful for detecting Hu-  
 CC Asp nucleic acids in in vitro assays and in Northern and Southern blots.  
 CC The present cDNA sequence encodes Human aspartyl protease 2b (Hu-Asp2b).  
 CC deltaTM protein which is obtained by the deletion of C-terminal  
 CC transmembrane and intracellular domains of Hu-Asp2b. Human Asp2b has beta-  
 CC secretase activity  
 XX  
 SQ Sequence 1287 BP; 271 A; 370 C; 384 G; 262 T; 0 U; 0 Other;

Alignment Scores:	Pred. No.:	Length:	1287
Score:	4,84e-14	Matches:	28
Percent Similarity:	144.00	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	100.00%	Indels:	0
DB:	4	Gaps:	0

US-10-726-967A-52 (1-28) x AAD13276 (1-1287)  
 QY 1 GLYVYRYYVVALGLMETHRYVALGLYSEKPROGQINHRLEUASNLLEUVALASP 20  
 DB 220 GGGTAACTACGTGAGATGACCGTGGGACAGCCCGCCGACAGCGCTCAACATCTCGTGGAT 279



Seq	Accession	Gene	Protein	Length	Score	Percent Similarity	Percent Identity	Local Similarity	Local Identity	Query Match	DB
1	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
2	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
3	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
4	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
5	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
6	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
7	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
8	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
9	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
10	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
11	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
12	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
13	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
14	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
15	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
16	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
17	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
18	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
19	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
20	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
21	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
22	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
23	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
24	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
25	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
26	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
27	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
28	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
29	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
30	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00	100.00	100.00	4	NCBI
31	U00001	Thy	Thy	1287	4.84e-14	144.00	100.00				

220 GGCTACTACGTGGAATGATACCGTGGGCGACCCCGCCGACGACGCTGCAACATCTCGTGGAT 21  
21 ThrGlySerSerAsnPhenAlaVal 28  
280 ACAGCGACGACGTACTTGTGCACTG 303

RESULT 14  
AAS11547  
ID AAS11547 standard; cDNA; 1287 BP.  
XX AAS11547;  
XX  
XX 24-OCT-2001 (first entry)  
XX  
XX Human cDNA encoding Human-pro-Asp 2 (b) delta TM.  
XX  
XX Human; Aspartyl protease; beta-secretase; neurotropic; ASP2;  
XX neuroprotective; amyloid protein precursor; APP; Alzheimer's disease;  
XX amyloid-beta; Abeta; Human-pro-Asp 2(b) delta TM; ss; mutant.  
XX  
XX Homo sapiens.  
XX Synthetic.  
XX  
XX Key Location/Qualifiers  
XX CDS 1..1287  
XX /tag=a  
XX /product="Human-pro-Asp 2 (b) delta TM"  
XX  
XX WO200149098-A2.  
XX  
XX 12-JUL-2001.  
XX  
XX 09-MAY-2001; 2001WO-IB000798.  
XX PF  
XX 09-MAY-2001; 2001WO-IB000798.  
XX PR  
XX  
XX (BIEN/) BIENKOWSKI M J.  
XX (GURN/) GURNEY M E.  
XX (HEIN/) HEINRIKSON R L.  
XX (PARO/) PARODI L A.  
XX (YANR/) YAN R.  
XX  
XX Blenkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;  
XX WPI; 2001-502549/55.  
XX  
XX Novel purified polypeptide comprising fragment of mammalian aspartyl  
XX protease 2, lacking Asp2 transmembrane domain and retaining beta  
XX secretase activity of Asp2 useful for identifying inhibitors of Asp2  
XX activity.  
XX  
XX Disclosure; Page 166-167; 185pp; English.

The invention relates to a purified polypeptide comprising a fragment of mammalian aspartyl protease (Asp2) protein which lacks the Asp2 transmembrane domain and the Asp2 protein, and where the polypeptide and the fragment retain the beta-secretase activity of the mammalian Asp2 protein. The invention also details polynucleotides for the Asp proteins and vectors expressing them, and a polypeptide (isoform of amyloid protein precursor (APP) comprising the amino acid sequence of an APP or its fragment containing an APP cleavage site recognizable by a mammalian beta-secretase, and further comprising two lysine residues at the carboxyl terminus of the amino acid sequence of the mammalian APP or APP fragment. Also included in the invention are methods of identifying modulators or inhibitors of Asp2. Modulators and inhibitors of Asp2 are useful for treating Alzheimer's disease. APP is useful in methods for identifying inhibitors or modulators of human Asp2 activity and amyloid-beta (Abeta) peptide production. APP is also useful in designing therapeutics for the treatment or prevention of Alzheimer's disease. APP comprising the APP-Sw-beta-secretase peptide sequence (NDA), which is associated with increased levels of Abeta processing is useful in assays relating the Alzheimer's disease. The expression vector is useful for recombinantly expressing APP. Nucleic acids that hybridise to Asp

CC oligonucleotides are useful as probes or primers. The probes are useful  
 CC for detecting Hu-Asp nucleic acids in in vitro assays and in Northern and  
 CC Southern blots. The present sequence encodes Human-pro- Asp 2 (b) delta TM  
 CC protein, which lacks the C-terminal transmembrane domain

XX SQ Sequence 1287 BP; 271 A; 370 C; 384 G; 262 T; 0 U; 0 Other;

#### Alignment Scores:

Score:	4.84e-14	Length:	1287
Percent Similarity:	100.00%	Matches:	28
Best Local Similarity:	100.00%	Conservative:	0
Query Match:	100.00%	Mismatches:	0
		Indels:	0
		Gaps:	0

US-10-726-967A-52 (1-28) x ABL52487 (1-1287)

OY 1 GYTYTYTYTYValGluMetThrValGlySerProProGlnThrLeuAenIleLeuValAsp 20

Db 220 GGCTACTACGTGAGATGACCGTGGGACGCCCGCAGACGCTCAACATCTGTGTGAT 279

OY 21 ThrGlySerSerAenPheAlaVal 28

Db 280 ACAGGACGACGTAACTTTCAGTGG 303

#### RESULT 15

ABL52487  
 ID ABL52487 standard; cDNA; 1287 BP.

XX AC ABL52487;

XX DT 16-JUN-2002 (first entry)

XX DE Human Asp-2(b)deltaTM nucleotide sequence SEQ ID NO:50.

XX KM Human; Asp-1; Asp-2; aspartyl protease; Alzheimer's disease; proteolytic;  
 XX chromosome 11q23.3-24.1; gene; ss.

XX OS Homo sapiens.

XX FT Key Location/Qualifiers

FT CDS 1..1287

FT /tag= a

FT /product= "Human Asp-2(b)delta TM"

XX GB2367060-A.

XX PD 27-MAR-2002.

XX PF 29-OCT-2001; 2001GB-00025934.

XX PR 23-SEP-1999; 99US-00404133.

XX PR 23-SEP-1999; 98US-0155493P.

XX PR 23-SEP-1999; 99MO-US020881.

XX PR 13-OCT-1999; 99US-00416901.

XX PR 06-DEC-1999; 99US-0169232P.

XX PR 22-SEP-2000; 2000GB-00023315.

XX PA (PHAA ) PHARMACIA & UPJOHN CO.

XX PI Bienkowskaki MJ, Gurney M;

XX DR WPI; 2002-397167/43.

XX DR P-PSDB; ABB78607.

XX PT Human aspartyl protease 1 substrates useful in assays to detect aspartyl  
 XX protease activity, e.g. for the diagnosis of Alzheimer's disease.  
 XX Example 10; Page 137; 182pp; English.  
 CC The present invention describes a human aspartyl protease 1 (hu-Asp1)  
 CC substrate (I) which comprises a peptide of no more than 50 amino acids,  
 CC and which comprises the 8 amino acid sequence Gly-Leu-Ala-Leu-Ala-Leu-

CC Glu-Pro. Also described are: (1) a method (II) for assaying hu-Asp1  
 CC proteolytic activity, comprising: (a) contacting a hu-Asp1 protein with  
 CC (I) under acidic conditions; and (b) determining the level of hu-Asp1  
 CC proteolytic activity; (2) a purified polynucleotide (III) comprising a  
 CC nucleotide sequence that hybridizes under stringent conditions to the non  
 CC -coding strand complementary to a defined 1804 nucleotide sequence (see  
 CC ABL52456) where the nucleotide sequence encodes a polypeptide having Asp1  
 CC proteolytic activity and lacks nucleotides encoding a transmembrane  
 CC domain; (3) a purified polynucleotide (III') comprising a sequence that  
 CC hybridizes under stringent conditions to (III) (the nucleotide sequence  
 CC encodes a polypeptide further lacking a pro-peptide domain corresponding  
 CC to amino acids 23-62 of hu-Asp1 (see ABB78589)); (4) a vector (IV)  
 CC comprising (III) or (III'); and (5) a host cell (V) transformed or  
 CC transfected with (III), (III') and/or (IV). The hu-Asp1 protease  
 CC substrate (I) may be used as an enzyme substrate in assays to detect  
 CC aspartyl protease activity, (II) and therefore diagnose diseases  
 CC associated with aberrant hu-Asp1 expression and activity such as  
 CC Alzheimer's disease. Hu-Asp1 has been localised to chromosome 21, while  
 CC hu-Asp2 has been localised to chromosome 11q23.3-24.1. The present  
 CC sequence encodes human Asp-2(b)deltaTM, which is given in an example from  
 CC the present invention

XX SQ Sequence 1287 BP; 271 A; 370 C; 384 G; 262 T; 0 U; 0 Other;

#### Alignment Scores:

Score:	4.84e-14	Length:	1287
Percent Similarity:	100.00%	Matches:	28
Best Local Similarity:	100.00%	Conservative:	0
Query Match:	100.00%	Mismatches:	0
		Indels:	0
		Gaps:	0

US-10-726-967A-52 (1-28) x ABL52487 (1-1287)

OY 1 GYTYTYTYTYValGluMetThrValGlySerProProGlnThrLeuAenIleLeuValAsp 20

Db 220 GGCTACTACGTGAGATGACCGTGGGACGCCCGCAGACGCTCAACATCTGTGTGAT 279

OY 21 ThrGlySerSerAenPheAlaVal 28

Db 280 ACAGGACGACGTAACTTTCAGTGG 303

Search completed: July 27, 2005, 17:25:06  
 Job time : 440 secs

GenCore version 5.1.6  
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OM protein - nucleic search, using frame\_plus.p2n model

Run on: July 27, 2005, 16:49:48 ; Search time 132 Seconds  
(without alignments)  
347.089 Million cell updates/sec

Title: US-10-726-967A-52

Perfect score: 144  
Sequence: 1 GYVEMTGSPPQTINILVDTGSSNPAV 28

Scoring table:  
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Ygapop 10.0 , Ygapext 0.5  
Fgapop 6.0 , Fgapext 7.0  
Delop 6.0 , Delext 7.0

Searched: 1202784 seqs, 818138359 residues

Total number of hits satisfying chosen parameters: 2405568

Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

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-DB=Issued\_Patents.NA -QFMT=fastap -SUFFIX=p2n.rn1 -MINMATCH=0.1 -LOOPEXT=0  
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-FGAPEXT=7 -YGAPOP=10 -YGAPEXT=0.5 -DELOP=6 -DELEXT=7

Database :

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3: /cgcn2.6/prodata/1/ina/6A.COMB.seq.\*  
4: /cgcn2.6/prodata/1/ina/6B.COMB.seq.\*  
5: /cgcn2.6/prodata/1/ina/PCTUS.COMB.seq.\*  
6: /cgcn2.6/prodata/1/ina/backfile1.seq.\*

Pred. No. is the number of results predicted by chance to have a  
score greater than or equal to the score of the result being printed,  
and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	144	100.0	1278	3	US-09-548-372D-27
2	144	100.0	1278	3	US-09-548-367D-27
3	144	100.0	1278	4	US-09-551-853D-27
4	144	100.0	1278	4	US-09-416-901B-50
5	144	100.0	1278	4	US-09-548-376D-27
6	144	100.0	1278	4	US-09-794-927A-27
7	144	100.0	1278	4	US-09-548-373D-27
8	144	100.0	1278	4	US-09-795-847B-27
9	144	100.0	1278	4	US-09-869-414-27
10	144	100.0	1278	4	US-09-548-366F-27
11	144	100.0	1278	4	US-09-548-368D-27
12	144	100.0	1278	4	US-09-794-925A-27

13	144	100.0	1278	4	US-09-806-194A-27	Sequence 27, Appl
14	144	100.0	1287	3	US-09-548-372D-50	Sequence 50, Appl
15	144	100.0	1287	3	US-09-548-367D-50	Sequence 50, Appl
16	144	100.0	1287	4	US-09-551-853D-50	Sequence 50, Appl
17	144	100.0	1287	4	US-09-416-901B-50	Sequence 50, Appl
18	144	100.0	1287	4	US-09-548-376D-50	Sequence 50, Appl
19	144	100.0	1287	4	US-09-794-927A-50	Sequence 50, Appl
20	144	100.0	1287	4	US-09-548-373D-50	Sequence 50, Appl
21	144	100.0	1287	4	US-09-795-847B-50	Sequence 50, Appl
22	144	100.0	1287	4	US-09-869-414-50	Sequence 50, Appl
23	144	100.0	1287	4	US-09-548-366F-50	Sequence 50, Appl
24	144	100.0	1287	4	US-09-548-368D-50	Sequence 50, Appl
25	144	100.0	1287	4	US-09-794-925A-50	Sequence 50, Appl
26	144	100.0	1302	3	US-09-548-372D-25	Sequence 25, Appl
27	144	100.0	1302	3	US-09-548-367D-25	Sequence 25, Appl
28	144	100.0	1302	4	US-09-551-853D-25	Sequence 25, Appl
29	144	100.0	1302	4	US-09-416-901B-25	Sequence 25, Appl
30	144	100.0	1302	4	US-09-548-376D-25	Sequence 25, Appl
31	144	100.0	1302	4	US-09-794-927A-25	Sequence 25, Appl
32	144	100.0	1302	4	US-09-548-373D-25	Sequence 25, Appl
33	144	100.0	1302	4	US-09-795-847B-25	Sequence 25, Appl
34	144	100.0	1302	4	US-09-869-414-25	Sequence 25, Appl
35	144	100.0	1302	4	US-09-548-366F-25	Sequence 25, Appl
36	144	100.0	1302	4	US-09-548-368D-25	Sequence 25, Appl
37	144	100.0	1302	4	US-09-794-925A-25	Sequence 25, Appl
38	144	100.0	1302	4	US-09-806-194A-25	Sequence 25, Appl
39	144	100.0	1305	3	US-09-548-372D-52	Sequence 52, Appl
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43	144	100.0	1305	4	US-09-548-376D-52	Sequence 52, Appl
44	144	100.0	1305	4	US-09-794-927A-52	Sequence 52, Appl
45	144	100.0	1305	4	US-09-548-373D-52	Sequence 52, Appl

#### ALIGNMENTS

RESULT 1  
US-09-548-372D-27  
Sequence 27, Application US/09548372D  
Patent No. 6420534  
GENERAL INFORMATION:  
APPLICANT: GURNEY ET AL.  
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES  
TITLE OF INVENTION: THEREOF  
FILE REFERENCE: 29915/62801  
CURRENT APPLICATION NUMBER: US/09/548, 372D  
CURRENT FILING DATE: 2000-04-12  
PRIOR APPLICATION NUMBER: US 60/155,493  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: US 09/404,133  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: PCT/US99/20881  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: US 60/101,594  
PRIOR FILING DATE: 1998-09-24  
NUMBER OF SEQ ID NOS: 73  
SOFTWARE: PatentIn version 3.1  
SEQ ID NO 27  
LENGTH: 1278  
TYPE: DNA  
ORGANISM: Homo sapiens  
US-09-548-372D-27  
Alignment Scores:  
Pred. No.: 4.05e-16  
Score: 144.00  
Percent Similarity: 100.00%  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
Matches: 1278  
Conservative: 28  
Mismatches: 0  
Indels: 0  
Gaps: 0  
US-10-726-967A-52 (1-28) x US-09-548-372D-27 (1-1278)

QY 1 G|Y|T|Y|T|Y|V|A|G|L|U|E|T|H|V|A|G|Y|S|E|R|P|R|O|G|I|N|T|H|L|E|U|A|N|I|L|E|U|V|A|L|A|S|P 20  
Db 136 GGCCTACTACGTGAGATGACCGTGGGCGAGCCCCCGGACAGCTCAACATCTGTGGAT 195  
QY 21 Thg|Y|S|E|S|E|A|P|H|E|A|L|A|V|A|L 28  
Db 196 ACAGGACAGCACTTAACCTTGCACTG 219

RESULT 2  
US-09-548-367D-27  
Sequence 27, Application US/09548367D  
Patent No. 6440698  
GENERAL INFORMATION:  
APPLICANT: GURNEY ET AL.  
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES  
FILE REFERENCE: 29915/6280H  
CURRENT APPLICATION NUMBER: US/09/548,367D  
CURRENT FILING DATE: 2000-04-12  
PRIOR APPLICATION NUMBER: US 60/155,493  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: US 09/404,133  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: PCT/US99/20881  
PRIOR FILING DATE: 1999-09-23  
NUMBER OF SEQ ID NOS: 73  
SOFTWARE: Patentin version 3.1  
SEQ ID NO 27  
LENGTH: 1278  
TYPE: DNA  
ORGANISM: Homo sapiens  
US-09-548-367D-27

Alignment Scores:  
Pred. No.: 4,05e-16 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
Gaps: 0

US-10-726-967A-52 (1-28) x US-09-548-367D-27 (1-1278)

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Db 136 GGCCTACTACGTGAGATGACCGTGGGCGAGCCCCCGGACAGCTCAACATCTGTGGAT 195

QY 21 Thg|Y|S|E|S|E|A|P|H|E|A|L|A|V|A|L 28  
Db 196 ACAGGACAGCACTTAACCTTGCACTG 219

RESULT 3  
US-09-551-853D-27  
Sequence 27, Application US/09551853D  
Patent No. 6500667  
GENERAL INFORMATION:  
APPLICANT: GURNEY ET AL.  
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES  
FILE REFERENCE: 29915/6280L  
CURRENT APPLICATION NUMBER: US/09/551,853D  
CURRENT FILING DATE: 2000-04-18  
PRIOR APPLICATION NUMBER: US 60/155,493  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: US 09/404,133  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: PCT/US99/20881  
PRIOR FILING DATE: 1999-09-23  
NUMBER OF SEQ ID NOS: 73  
SOFTWARE: Patentin version 3.1  
SEQ ID NO 27  
LENGTH: 1278  
TYPE: DNA  
ORGANISM: Homo sapiens  
US-09-551-853D-27

NUMBER OF SEQ ID NOS: 73  
SOFTWARE: Patentin version 3.1  
SEQ ID NO 27  
LENGTH: 1278  
TYPE: DNA  
ORGANISM: Homo sapiens  
US-09-551-853D-27

Alignment Scores:  
Pred. No.: 4,05e-16 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
Gaps: 0

US-10-726-967A-52 (1-28) x US-09-551-853D-27 (1-1278)

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Db 136 GGCCTACTACGTGAGATGACCGTGGGCGAGCCCCCGGACAGCTCAACATCTGTGGAT 195

QY 21 Thg|Y|S|E|S|E|A|P|H|E|A|L|A|V|A|L 28  
Db 196 ACAGGACAGCACTTAACCTTGCACTG 219

RESULT 4  
US-09-416-901B-27  
Sequence 27, Application US/09416901B  
Patent No. 6699671  
GENERAL INFORMATION:  
APPLICANT: GURNEY ET AL.  
TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES  
FILE REFERENCE: 29915/6280A  
CURRENT APPLICATION NUMBER: US/09/416,901B  
CURRENT FILING DATE: 1999-10-13  
PRIOR APPLICATION NUMBER: US 60/155,493  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: US 09/404,133  
PRIOR FILING DATE: 1999-09-23  
PRIOR APPLICATION NUMBER: PCT/US99/20881  
PRIOR FILING DATE: 1999-09-23  
NUMBER OF SEQ ID NOS: 72  
SOFTWARE: Patentin version 3.1  
SEQ ID NO 27  
LENGTH: 1278  
TYPE: DNA  
ORGANISM: Homo sapiens  
US-09-416-901B-27

Alignment Scores:  
Pred. No.: 4,05e-16 Length: 1278  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
Gaps: 0

US-10-726-967A-52 (1-28) x US-09-416-901B-27 (1-1278)

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Db 136 GGCCTACTACGTGAGATGACCGTGGGCGAGCCCCCGGACAGCTCAACATCTGTGGAT 195

QY 21 Thg|Y|S|E|S|E|A|P|H|E|A|L|A|V|A|L 28  
Db 196 ACAGGACAGCACTTAACCTTGCACTG 219

RESULT 5  
US-09-548-376D-27

US-09-134-92/A-21

APPLICANT: PARODI, LUISE

Alignment Scores:	
Pred. No.:	4.05e-16
Score:	144.00
Length:	1278
Matches:	28

PRIOR FILING DATE: 1999-09-23

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; PRIOR APPLICATION NUMBER: US 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: PatentIn version 3.1
; SEQ ID NO 27
; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-09-548-368D-27

Alignment Scores:
Pred. No.: 4,05e-16 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 4 Gaps: 0

US-10-726-967a-52 (1-28) x US-09-548-368D-27 (1-1278)
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DB 136 G|G|C|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|T|G|G|C|A|G|C|C|C|C|C|G|C|A|G|C|T|C|A|C|A|C|T|C|T|G|T|G|A|T 195

QY 21 T|H|G|Y|S|E|S|E|A|N|P|H|E|A|L|A|V|A|I 28
DB 196 A|C|A|G|G|A|G|C|A|G|T|A|C|T|T|T|G|C|A|G|T|G 219

RESULT 12
US-09-794-925A-27
; Sequence 27, Application US/09794925A
; Patent No. 6828117
; GENERAL INFORMATION:
; APPLICANT: Gurney et al.
; TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses
; FILE REFERENCE: 29915/6280H1
; CURRENT APPLICATION NUMBER: US/09/794,925A
; CURRENT FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 74
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 27
; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-09-794-925A-27

Alignment Scores:
Pred. No.: 4,05e-16 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 4 Gaps: 0

US-10-726-967a-52 (1-28) x US-09-794-925A-27 (1-1278)
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DB 136 G|G|C|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|T|G|G|C|A|G|C|C|C|C|C|G|C|A|G|C|T|C|A|C|A|C|T|C|T|G|T|G|A|T 195

QY 21 T|H|G|Y|S|E|S|E|A|N|P|H|E|A|L|A|V|A|I 28
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DB 136 G|G|C|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|T|G|G|C|A|G|C|C|C|C|C|G|C|A|G|C|T|C|A|C|A|C|T|C|T|G|T|G|A|T 195
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DB 196 A|C|A|G|G|A|G|C|A|G|T|A|C|T|T|T|G|C|A|G|T|G 219

RESULT 13
US-09-806-194A-27
; Sequence 27, Application US/09806194A
; Patent No. 6835565
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrikson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; APPLICANT: Pharmacia & Upjohn Company
; TITLE OF INVENTION: Alzheimer's Disease Secretase
; FILE REFERENCE: 6177.P CP
; CURRENT APPLICATION NUMBER: US/09/806,194A
; CURRENT FILING DATE: 2001-09-17
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 27
; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-09-806-194A-27

Alignment Scores:
Pred. No.: 4,05e-16 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 4 Gaps: 0

US-10-726-967a-52 (1-28) x US-09-806-194A-27 (1-1278)
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QY 21 T|H|G|Y|S|E|S|E|A|N|P|H|E|A|L|A|V|A|I 28
DB 196 A|C|A|G|G|A|G|C|A|G|T|A|C|T|T|T|G|C|A|G|T|G 219

RESULT 14
US-09-548-372D-50
; Sequence 50, Application US/09548372D
; Patent No. 6420534
; GENERAL INFORMATION:
; APPLICANT: GURNEY ET AL.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES
; FILE REFERENCE: 29915/62801
; CURRENT APPLICATION NUMBER: US/09/548,372D
; CURRENT FILING DATE: 2000-04-12
; PRIOR APPLICATION NUMBER: US 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: PatentIn version 3.1
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
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; ORGANISM: Artificial sequence
; FEATURE:
; OTHER INFORMATION: Hu-Asp2 (b) delta TM
US-09-548-372D-50

Alignment Scores:
Pred. No.:      4.09e-16      Length:      1287
Score:          144.00      Matches:      28
Percent Similarity: 100.00%      Conservative: 0
Best Local Similarity: 100.00%      Mismatches: 0
Query Match:      100.00%      Indels:      0
DB:              3          Gaps:      0

US-10-726-967A-52 (1-28) x US-09-548-372D-50 (1-1287)

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QY      21  T h r G l y S e r S e r A s n P h e A l a V a l 28
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RESULT 15
US-09-548-367D-50
; Sequence 50, Application US/09548367D
; Patent No. 6440698
; GENERAL INFORMATION:
; APPLICANT: GURNEY ET AL.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR AND USES
; FILE OF INVENTION: THEREOF
; FILE REFERENCE: 29915/6280H
; CURRENT APPLICATION NUMBER: US/09/548,367D
; CURRENT FILING DATE: 2000-04-12
; PRIOR APPLICATION NUMBER: US 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20861
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: US 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: PatentIn version 3.1
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial sequence
; FEATURE:
; OTHER INFORMATION: Hu-Asp2 (b) delta TM
US-09-548-367D-50

Alignment Scores:
Pred. No.:      4.09e-16      Length:      1287
Score:          144.00      Matches:      28
Percent Similarity: 100.00%      Conservative: 0
Best Local Similarity: 100.00%      Mismatches: 0
Query Match:      100.00%      Indels:      0
DB:              3          Gaps:      0

US-10-726-967A-52 (1-28) x US-09-548-367D-50 (1-1287)

QY      1  G I Y T Y T Y R V A l G l u M e t h r V a l G l y S e r P r o P r o G l n T h l e u A s n l l e l e u V a l A s p 20
      |||
Db      220 G G C T A C T A C G T G A G A T G A C C G T G G C A G C C C C C G C A G C G T C A C A C A T C C T G G T G A T 279

QY      21  T h r G l y S e r S e r A s n P h e A l a V a l 28
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Db      280 A C A G G C A G C A G T A C T T T G C A G T G 303

Search completed: July 27, 2005, 18:52:41
Job time : 134 secs
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GenCore version 5.1.6  
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OM protein - nucleic search, using frame\_plus.p2n model

Run on: July 27, 2005, 17:18:01 ; Search time 627 Seconds  
(without alignments)  
288.765 Million cell updates/sec

Title: US-10-726-967A-52  
Perfect score: 144  
Sequence: 1 GYVEMTVGSPPTQTLIVDTGSSNFAV 28

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Ygapop 10.0 , Ygapext 0.5  
Fgapop 6.0 , Fgapext 7.0  
Delop 6.0 , Delext 7.0

Searched: 7277826 seqs, 3233139505 residues  
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Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

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21: /cgn2\_6/ptodata/1/pubpna/US10I\_PUBCOMB.seq:\*  
22: /cgn2\_6/ptodata/1/pubpna/US11A\_PUBCOMB.seq:\*  
23: /cgn2\_6/ptodata/1/pubpna/US11\_NEW\_PUB.seq:\*  
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Pred. No. is the number of results predicted by chance to have a

score greater than or equal to the score of the result being printed,  
and is derived by analysis of the total score distribution.

SUMMARIES

Result No.	Score	Query Match	Query Length	DB	ID	Description
1	144	100.0	1278	9	US-09-794-927-27	Sequence 27, Appl
2	144	100.0	1278	9	US-09-795-847-27	Sequence 27, Appl
3	144	100.0	1278	9	US-09-794-743-27	Sequence 27, Appl
4	144	100.0	1278	9	US-09-794-748-27	Sequence 27, Appl
5	144	100.0	1278	9	US-09-794-925-27	Sequence 27, Appl
6	144	100.0	1278	9	US-09-681-442-27	Sequence 27, Appl
7	144	100.0	1278	10	US-09-681-442-27	Sequence 27, Appl
8	144	100.0	1278	10	US-09-548-366-27	Sequence 27, Appl
9	144	100.0	1278	18	US-10-652-927-27	Sequence 27, Appl
10	144	100.0	1278	18	US-10-652-830-27	Sequence 27, Appl
11	144	100.0	1278	19	US-10-652-045-27	Sequence 27, Appl
12	144	100.0	1278	20	US-10-476-935-27	Sequence 27, Appl
13	144	100.0	1278	21	US-10-940-867-27	Sequence 27, Appl
14	144	100.0	1278	21	US-10-477-076-27	Sequence 27, Appl
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17	144	100.0	1287	9	US-09-794-743-50	Sequence 50, Appl
18	144	100.0	1287	9	US-09-794-748-50	Sequence 50, Appl
19	144	100.0	1287	9	US-09-794-925-50	Sequence 50, Appl
20	144	100.0	1287	9	US-09-681-442-50	Sequence 50, Appl
21	144	100.0	1287	10	US-09-869-414-50	Sequence 50, Appl
22	144	100.0	1287	10	US-09-548-366-50	Sequence 50, Appl
23	144	100.0	1287	18	US-10-652-927-50	Sequence 50, Appl
24	144	100.0	1287	18	US-10-652-830-50	Sequence 50, Appl
25	144	100.0	1287	19	US-10-652-045-50	Sequence 50, Appl
26	144	100.0	1287	20	US-10-476-935-50	Sequence 50, Appl
27	144	100.0	1287	21	US-10-940-867-50	Sequence 50, Appl
28	144	100.0	1302	9	US-09-794-927-25	Sequence 25, Appl
29	144	100.0	1302	9	US-09-795-847-25	Sequence 25, Appl
30	144	100.0	1302	9	US-09-794-743-25	Sequence 25, Appl
31	144	100.0	1302	9	US-09-794-748-25	Sequence 25, Appl
32	144	100.0	1302	9	US-09-794-925-25	Sequence 25, Appl
33	144	100.0	1302	9	US-09-681-442-25	Sequence 25, Appl
34	144	100.0	1302	10	US-09-869-414-25	Sequence 25, Appl
35	144	100.0	1302	10	US-09-548-366-25	Sequence 25, Appl
36	144	100.0	1302	18	US-10-652-927-25	Sequence 25, Appl
37	144	100.0	1302	18	US-10-652-830-25	Sequence 25, Appl
38	144	100.0	1302	19	US-10-652-045-25	Sequence 25, Appl
39	144	100.0	1302	20	US-10-476-935-25	Sequence 25, Appl
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41	144	100.0	1302	21	US-10-477-076-25	Sequence 25, Appl
42	144	100.0	1305	9	US-09-794-927-52	Sequence 52, Appl
43	144	100.0	1305	9	US-09-795-847-52	Sequence 52, Appl
44	144	100.0	1305	9	US-09-794-743-52	Sequence 52, Appl
45	144	100.0	1305	9	US-09-794-748-52	Sequence 52, Appl

ALIGNMENTS

RESULT 1  
US-09-794-927-27  
; Sequence 27, Application US/09794927  
; Patent No. US20010016324A1  
; GENERAL INFORMATION:  
; APPLICANT: Gurney, Mark E.  
; APPLICANT: Bienkowski, Michael J.  
; APPLICANT: Heinrikson, Robert L.  
; APPLICANT: Parodi, Luis A.  
; APPLICANT: Yan, Ridiang  
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND  
; TITLE OF INVENTION: THEREFOR  
; FILE REFERENCE: 28341/6280FG  
; CURRENT APPLICATION NUMBER: US/09/794, 927  
; CURRENT FILING DATE: 2001-02-27

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/ PRIOR APPLICATION NUMBER: 09/416,901
/ PRIOR FILING DATE: 1999-10-13
/ PRIOR APPLICATION NUMBER: 60/155,493
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: 09/404,133
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: PCT/US99/20881
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: 60/101,594
/ PRIOR FILING DATE: 1998-09-24
/ NUMBER OF SEQ ID NOS: 73
/ SOFTWARE: Patentln Ver. 2.0
/ SEQ ID NO 27
/ LENGTH: 1278
/ TYPE: DNA
/ ORGANISM: Homo sapiens
US-09-794-927-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0

US-10-726-967a-52 (1-28) x US-09-794-927-27 (1-1278)

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Db 136 G|G|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|T|G|G|C|A|G|C|C|C|C|C|G|C|A|G|C|T|C|A|C|T|C|T|G|T|G|A|T 195

QY 21 T|H|G|I|S|E|S|E|S|E|A|N|P|H|E|A|I|A|V|A|I 28
Db 196 A|C|A|G|G|C|A|G|C|A|G|T|A|C|T|T|G|C|A|G|T|G 219

RESULT 2
US-09-795-847-27
/ Sequence 27, Application US/09795847
/ Patent No. US20010018208A1
/ GENERAL INFORMATION:
/ APPLICANT: Gurney, Mark E.
/ APPLICANT: Bienkowski, Michael J.
/ APPLICANT: Heinrichson, Robert L.
/ APPLICANT: Parodi, Luis A.
/ APPLICANT: Yan, Riqiang
/ TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND
/ TITLE OF INVENTION: USES
/ FILE REFERENCE: 28341/6280DE
/ CURRENT APPLICATION NUMBER: US/09/795,847
/ CURRENT FILING DATE: 2001-02-28
/ PRIOR APPLICATION NUMBER: 09/416,901
/ PRIOR FILING DATE: 1999-10-13
/ PRIOR APPLICATION NUMBER: 60/155,493
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: 09/404,133
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: PCT/US99/20881
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: 60/101,594
/ PRIOR FILING DATE: 1998-09-24
/ NUMBER OF SEQ ID NOS: 73
/ SOFTWARE: Patentln Ver. 2.0
/ SEQ ID NO 27
/ LENGTH: 1278
/ TYPE: DNA
/ ORGANISM: Homo sapiens
US-09-795-847-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0
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Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0

US-10-726-967a-52 (1-28) x US-09-795-847-27 (1-1278)

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QY 21 T|H|G|I|S|E|S|E|S|E|A|N|P|H|E|A|I|A|V|A|I 28
Db 196 A|C|A|G|G|C|A|G|C|A|G|T|A|C|T|T|G|C|A|G|T|G 219

RESULT 3
US-09-794-743-27
/ Sequence 27, Application US/09794743
/ Patent No. US20010021391A1
/ GENERAL INFORMATION:
/ APPLICANT: Gurney, Mark E.
/ APPLICANT: Bienkowski, Michael J.
/ APPLICANT: Heinrichson, Robert L.
/ APPLICANT: Parodi, Luis A.
/ APPLICANT: Yan, Riqiang
/ TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND
/ TITLE OF INVENTION: USES
/ FILE REFERENCE: 28341/6280BC
/ CURRENT APPLICATION NUMBER: US/09/794,743
/ CURRENT FILING DATE: 2001-02-27
/ PRIOR APPLICATION NUMBER: 09/416,901
/ PRIOR FILING DATE: 1999-10-13
/ PRIOR APPLICATION NUMBER: 60/155,493
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: 09/404,133
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: PCT/US99/20881
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: 60/101,594
/ PRIOR FILING DATE: 1998-09-24
/ NUMBER OF SEQ ID NOS: 73
/ SOFTWARE: Patentln Ver. 2.0
/ SEQ ID NO 27
/ LENGTH: 1278
/ TYPE: DNA
/ ORGANISM: Homo sapiens
US-09-794-743-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0

US-10-726-967a-52 (1-28) x US-09-794-743-27 (1-1278)

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QY 21 T|H|G|I|S|E|S|E|S|E|A|N|P|H|E|A|I|A|V|A|I 28
Db 196 A|C|A|G|G|C|A|G|C|A|G|T|A|C|T|T|G|C|A|G|T|G 219

RESULT 4
US-09-794-748-27
/ Sequence 27, Application US/09794748
/ Patent No. US20020037315A1
/ GENERAL INFORMATION:
/ APPLICANT: Gurney, Mark E.
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; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-09-794-925-27

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Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0

US-10-726-967A-52 (1-28) x US-09-794-925-27 (1-1278)

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21 ThrGlySeRSeRAnPheAlaVal 28
Db 196 ACAGGACAGATTAATTTCAGTG 219

RESULT 6
US-09-681-442-27
; Sequence 27, Application US/09681442
; Patent No. US20020081634A1
; GENERAL INFORMATION:
; APPLICANT: Guiney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrichson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; FILE REFERENCE: 28341/6280FG
; CURRENT APPLICATION NUMBER: US/09/681,442
; CURRENT FILING DATE: 2001-04-05
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 27
; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-09-681-442-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0

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QY 1 GYTYTYTYVAIGlUmEThrVAIGlYSeRPrOPrOGInThrLeuSAnIlleuVAlaAp 20
Db 136 GGCTACTACGGAGATGACCGTGGCGAGCCCCCGCAGACGCTCAACATCCTGGTGAT 195
21 ThrGlySeRSeRAnPheAlaVal 28

```

```

Db          196 ACAGCAGCAGTAAGTCTTGCACTG 219

RESULT 7
US-09-869-414-27
/ Sequence 27, Application US/09869414
/ Publication No. US20030077226A1
/ GENERAL INFORMATION:
/ APPLICANT: Beinikowski et al.
/ TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES THEREFOR
/ FILE REFERENCE: 28341/6280M
/ CURRENT APPLICATION NUMBER: US/09/869,414
/ PRIOR FILING DATE: 2001-06-27
/ PRIOR APPLICATION NUMBER: 09/416,901
/ PRIOR FILING DATE: 1999-10-13
/ PRIOR APPLICATION NUMBER: 60/155,493
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: 09/404,133
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: PCT/US99/20861
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: 60/101,594
/ PRIOR FILING DATE: 1998-09-24
/ NUMBER OF SEQ ID NOS: 73
/ SOFTWARE: PatentIn Ver. 2.0
/ SEQ ID NO 27
/ LENGTH: 1278
/ TYPE: DNA
/ ORGANISM: Homo sapiens
US-09-869-414-27

Alignment Scores:
Pred. No.:          2,82e-14          Length:          1278
Score:              144.00           Matches:          28
Percent Similarity: 100.00%           Conservative:    0
Best Local Similarity: 100.00%         Mismatches:     0
Query Match:        100.00%           Indels:          0
DB:                 10                Gaps:            0

US-10-726-967A-52 (1-28) x US-09-869-414-27 (1-1278)

QY          1  GIGYTYTTRVAGIGUMETHVAGIGYSETPROPGINTHRLAENILLEUVALASP 20
              |||||
Db          136 GGGCTACTACGCGGAGATGACCGGTGGGAGCCGCCCGCAGAGCTCAACATCCTGGTGAT 195
              |||||

QY          21  ThIGlySeSeRaaPneALaVal 28
              |||||
Db          196 ACAGCAGCAGTAAGTCTTGCACTG 219

RESULT 8
US-09-548-366-27
/ Sequence 27, Application US/09548366
/ Publication No. US20030104365A1
/ GENERAL INFORMATION:
/ APPLICANT: Gurney, Mark E.
/ APPLICANT: Bienkowski, Michael J.
/ APPLICANT: Heinrichson, Robert L.
/ APPLICANT: Parodi, Luis A.
/ APPLICANT: Yan, Riqiang
/ TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES THEREFOR
/ FILE REFERENCE: 28341/6280A
/ CURRENT APPLICATION NUMBER: US/09/548,366
/ PRIOR FILING DATE: 2000-04-12
/ PRIOR APPLICATION NUMBER: 60/155,493
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: 09/404,133
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: PCT/US99/20861
/ PRIOR FILING DATE: 1999-09-23
/ PRIOR APPLICATION NUMBER: 60/101,594
/ PRIOR FILING DATE: 1998-09-24
/ NUMBER OF SEQ ID NOS: 65

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; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 27
; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-09-548-366-27

Alignment Scores:
Pred. No.:      2,82e-14      Length:      1278
Score:          144.00      Matches:      28
Percent Similarity: 100.00%      Conservative: 0
Best Local Similarity: 100.00%      Mismatches:  0
Query Match:     100.00%      Indels:      0
DB:              18      Gaps:      0

US-10-726-967A-52 (1-28) x US-09-548-366-27 (1-1278)

QY      1 G|Y|T|Y|T|Y|V|a|G|U|Me|T|Th|V|a|G|Y|Se|P|Ro|G|In|Th|r|Leu|A|en|I|Leu|V|a|L|a|a|P 20
Db      136 GGCCTACTACGTGAGATATACCGTGGGACGCCCGGACAGACGCTCAACATCTCTGTGGAT 195

QY      21 Th|G|Y|Se|Se|A|en|P|he|Al|a|V|a|L 28
Db      196 ACAGGACAGCAGTAATTGACAGT 219

RESULT 9
US-10-652-927-27
; Sequence 27, Application US/10652927
; Publication No. US20040043408A1
; GENERAL INFORMATION:
; APPLICANT: Gurney et al.
; TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses
; FILE REFERENCE: 23915/6280N3
; CURRENT APPLICATION NUMBER: US/10/652,927
; CURRENT FILING DATE: 2003-08-29
; PRIOR APPLICATION NUMBER: 09/794,925
; PRIOR FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 74
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 27
; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-10-652-927-27

Alignment Scores:
Pred. No.:      2,82e-14      Length:      1278
Score:          144.00      Matches:      28
Percent Similarity: 100.00%      Conservative: 0
Best Local Similarity: 100.00%      Mismatches:  0
Query Match:     100.00%      Indels:      0
DB:              18      Gaps:      0

US-10-726-967A-52 (1-28) x US-10-652-927-27 (1-1278)

QY      1 G|Y|T|Y|T|Y|V|a|G|U|Me|T|Th|V|a|G|Y|Se|P|Ro|G|In|Th|r|Leu|A|en|I|Leu|V|a|L|a|a|P 20
Db      136 GGCCTACTACGTGAGATATACCGTGGGACGCCCGGACAGACGCTCAACATCTCTGTGGAT 195

QY      21 Th|G|Y|Se|Se|A|en|P|he|Al|a|V|a|L 28
Db      196 ACAGGACAGCAGTAATTGACAGT 219

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RESULT 10
US-10-652-830-27
; Sequence 27, Application US/10652830
; Publication No. US20040048303A1
; GENERAL INFORMATION:
; APPLICANT: Gurney et al.
; TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses
; FILE REFERENCE: 29915/6280N1
; CURRENT APPLICATION NUMBER: US/10/652,830
; CURRENT FILING DATE: 2003-08-29
; PRIOR APPLICATION NUMBER: 09/794,925
; PRIOR FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 74
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 27
; TYPE: DNA
; ORGANISM: Homo sapiens
US-10-652-830-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 18 Gaps: 0

US-10-726-967a-52 (1-28) x US-10-652-830-27 (1-1278)

QY 1 G1YTYTYrValG1uWetThrValG1ySerProGlnThrLeuAn11eLeuValAsp 20
Db 136 GGCTACTACGTGAGATGACCGTGGCGAGCCCCCGCAGAGCTCAACATCTCTGTGGAT 195
QY 21 ThG1ySerSerAnPheAlaVal 28
Db 196 ACAGGCAGCAGTAACCTTGCAAGTG 219

RESULT 11
US-10-652-045-27
; Sequence 27, Application US/10652045
; Publication No. US20040166507A1
; GENERAL INFORMATION:
; APPLICANT: Gurney et al.
; TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses
; FILE REFERENCE: 29915/6280N2
; CURRENT APPLICATION NUMBER: US/10/652,045
; CURRENT FILING DATE: 2003-08-29
; PRIOR APPLICATION NUMBER: 09/794,925
; PRIOR FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1998-09-23
; NUMBER OF SEQ ID NOS: 60/101,594
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 27
; TYPE: DNA
; ORGANISM: Homo sapiens
US-10-652-045-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 18 Gaps: 0

US-10-726-967a-52 (1-28) x US-10-652-045-27 (1-1278)

QY 1 G1YTYTYrValG1uWetThrValG1ySerProGlnThrLeuAn11eLeuValAsp 20
Db 136 GGCTACTACGTGAGATGACCGTGGCGAGCCCCCGCAGAGCTCAACATCTCTGTGGAT 195
QY 21 ThG1ySerSerAnPheAlaVal 28
Db 196 ACAGGCAGCAGTAACCTTGCAAGTG 219

RESULT 10
US-10-652-830-27
; Sequence 27, Application US/10652830
; Publication No. US20040048303A1
; GENERAL INFORMATION:
; APPLICANT: Gurney et al.
; TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses
; FILE REFERENCE: 29915/6280N1
; CURRENT APPLICATION NUMBER: US/10/652,830
; CURRENT FILING DATE: 2003-08-29
; PRIOR APPLICATION NUMBER: 09/794,925
; PRIOR FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 74
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 27
; TYPE: DNA
; ORGANISM: Homo sapiens
US-10-652-830-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 18 Gaps: 0

US-10-726-967a-52 (1-28) x US-10-652-830-27 (1-1278)

QY 1 G1YTYTYrValG1uWetThrValG1ySerProGlnThrLeuAn11eLeuValAsp 20
Db 136 GGCTACTACGTGAGATGACCGTGGCGAGCCCCCGCAGAGCTCAACATCTCTGTGGAT 195
QY 21 ThG1ySerSerAnPheAlaVal 28
Db 196 ACAGGCAGCAGTAACCTTGCAAGTG 219

RESULT 11
US-10-652-045-27
; Sequence 27, Application US/10652045
; Publication No. US20040166507A1
; GENERAL INFORMATION:
; APPLICANT: Gurney et al.
; TITLE OF INVENTION: Alzheimer's Disease Secretase, APP Substrates Therefor and Uses
; FILE REFERENCE: 29915/6280N2
; CURRENT APPLICATION NUMBER: US/10/652,045
; CURRENT FILING DATE: 2003-08-29
; PRIOR APPLICATION NUMBER: 09/794,925
; PRIOR FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1998-09-23
; NUMBER OF SEQ ID NOS: 60/101,594
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 27
; TYPE: DNA
; ORGANISM: Homo sapiens
US-10-652-045-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 18 Gaps: 0

US-10-726-967a-52 (1-28) x US-10-652-045-27 (1-1278)

QY 1 G1YTYTYrValG1uWetThrValG1ySerProGlnThrLeuAn11eLeuValAsp 20
Db 136 GGCTACTACGTGAGATGACCGTGGCGAGCCCCCGCAGAGCTCAACATCTCTGTGGAT 195
QY 21 ThG1ySerSerAnPheAlaVal 28
Db 196 ACAGGCAGCAGTAACCTTGCAAGTG 219
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; NUMBER OF SEQ ID NOS: 74
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 27
; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-10-652-045-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 19 Gaps: 0

US-10-726-967a-52 (1-28) x US-10-652-045-27 (1-1278)

QY 1 G1YTYTYrValG1uWetThrValG1ySerProGlnThrLeuAn11eLeuValAsp 20
Db 136 GGCTACTACGTGAGATGACCGTGGCGAGCCCCCGCAGAGCTCAACATCTCTGTGGAT 195
QY 21 ThG1ySerSerAnPheAlaVal 28
Db 196 ACAGGCAGCAGTAACCTTGCAAGTG 219

RESULT 12
US-10-476-935-27
; Sequence 27, Application US/10476935
; Publication No. US20040234976A1
; GENERAL INFORMATION:
; APPLICANT: Belinkowski et al.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; FILE REFERENCE: 28341/6280N1
; CURRENT APPLICATION NUMBER: US/10/476,935
; CURRENT FILING DATE: 2003-11-06
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: Patentin Ver. 2.0
; SEQ ID NO 27
; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-10-476-935-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 20 Gaps: 0

US-10-726-967a-52 (1-28) x US-10-476-935-27 (1-1278)

QY 1 G1YTYTYrValG1uWetThrValG1ySerProGlnThrLeuAn11eLeuValAsp 20
Db 136 GGCTACTACGTGAGATGACCGTGGCGAGCCCCCGCAGAGCTCAACATCTCTGTGGAT 195
QY 21 ThG1ySerSerAnPheAlaVal 28
Db 196 ACAGGCAGCAGTAACCTTGCAAGTG 219
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RESULT 13
US-10-940-867-27
; Sequence 27, Application US/10940867
; Publication No. US20050026256A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrichson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; APPLICANT: Pharmacia & Upjohn Company
; TITLE OF INVENTION: Alzheimer's Disease Secretase
; FILE REFERENCE: 6177.PCPA
; CURRENT APPLICATION NUMBER: US/10/940,867
; PRIOR FILING DATE: 2004-09-14
; PRIOR APPLICATION NUMBER: US 09/806,194
; PRIOR FILING DATE: 2001-03-26
; PRIOR APPLICATION NUMBER: US 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 49
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 27
; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-10-940-867-27

Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 21 Gaps: 0

US-10-726-967A-52 (1-28) x US-10-940-867-27 (1-1278)
QY 1 GATYTYTVVAlGUmEThrVAlGlySerProPrgInThrLeuAnllLeuValAsp 20
Db 136 GGCTACTAGTGAGATATACCGTGGCAGCCCCCGCAGACCTCAACATCTCTGTGGAT 195
QY 21 ThrglySerSerAsnPhelAlaVal 28
Db 196 ACAGGCAGCAGTAATTTCAGTG 219

RESULT 14
US-10-477-076-27
; Sequence 27, Application US/10477076
; Publication No. US20050080232A1
; GENERAL INFORMATION:
; APPLICANT: Beinkowski et al.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
; FILE REFERENCE: 28341/6280M2
; CURRENT APPLICATION NUMBER: US/10/477,076
; PRIOR FILING DATE: 2003-11-06
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 27
; LENGTH: 1278
; TYPE: DNA
; ORGANISM: Homo sapiens
US-10-477-076-27
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```
Alignment Scores:
Pred. No.: 2,82e-14 Length: 1278
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 21 Gaps: 0

US-10-726-967A-52 (1-28) x US-09-794-927-50 (1-1287)
QY 1 GATYTYTVVAlGUmEThrVAlGlySerProPrgInThrLeuAnllLeuValAsp 20
Db 220 GGCTACTAGTGAGATATACCGTGGCAGCCCCCGCAGACCTCAACATCTCTGTGGAT 279
QY 21 ThrglySerSerAsnPhelAlaVal 28
Db 280 ACAGGCAGCAGTAATTTCAGTG 303

RESULT 15
US-09-794-927-50
; Sequence 50, Application US/09794927
; Patent No. US20010016324A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
; APPLICANT: Bienkowski, Michael J.
; APPLICANT: Heinrichson, Robert L.
; APPLICANT: Parodi, Luis A.
; APPLICANT: Yan, Riqiang
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND
; TITLE OF INVENTION: USES
; FILE REFERENCE: 28341/6280M2
; CURRENT APPLICATION NUMBER: US/09/794,927
; PRIOR FILING DATE: 2001-02-27
; PRIOR APPLICATION NUMBER: 09/416,901
; PRIOR FILING DATE: 1999-10-13
; PRIOR APPLICATION NUMBER: 60/155,493
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 09/404,133
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: PCT/US99/20881
; PRIOR FILING DATE: 1999-09-23
; PRIOR APPLICATION NUMBER: 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 50
; LENGTH: 1287
; TYPE: DNA
; ORGANISM: Artificial Sequence
; FEATURE:
; OTHER INFORMATION: Description of Artificial Sequence: Hu-Asp2 (b)
US-09-794-927-50

Alignment Scores:
Pred. No.: 2,84e-14 Length: 1287
Score: 144.00 Matches: 28
Percent Similarity: 100.00% Conservative: 0
Best Local Similarity: 100.00% Mismatches: 0
Query Match: 100.00% Indels: 0
DB: 9 Gaps: 0
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Job time : 629 secs

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OM protein - nucleic search, using frame\_plus\_p2n model

Run on: July 27, 2005, 13:56:30 ; Search time 3149 Seconds  
(without alignments)  
338.457 Million cell updates/sec

Title: US-10-726-967A-52

Perfect score: 144  
Sequence: 1 GYVEMTVGSPPTQINLVDTGSSNPAV 28

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Ygapop 10.0 , Ygapext 0.5  
Fgapop 6.0 , Fgapext 7.0  
Delop 6.0 , Delext 7.0

Searched: 34239544 seqs, 19032134700 residues

Total number of hits satisfying chosen parameters: 68479088

Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

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-DB=EST -OPMT=fastlap -SUFFIX=p2n.rst -MINMATCH=0.1 -LOOPEXT=0 -LOOPEXT=0  
-UNITS=bits -START=1 -END=1 -MATRIX=blosum62 -TRANS=human40.cdi -LIST=45  
-DOCALLIGN=200 -THR\_SCORE=pct -THR\_MAX=100 -THR\_MIN=0 -ALIGN=15 -MODE=LOCAL  
-OUTFMT=PCO -NORM=ext -HEAPSIZE=500 -MINLEN=0 -MAXLEN=200000000  
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-DEV TIMEOUT=120 -WARN TIMEOUT=30 -THREADS=1 -XGAPOP=10 -XGAPEXT=0.5 -FGAPOP=6  
-FAPEXT=7 -YGAPOP=10 -YGAPEXT=0.5 -DELOP=6 -DELEXT=7

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EST:  
1: gb\_est1:\*  
2: gb\_est2:\*  
3: gb\_hic:\*  
4: gb\_est3:\*  
5: gb\_est4:\*  
6: gb\_est5:\*  
7: gb\_est6:\*  
8: gb\_gsa1:\*  
9: gb\_gsa2:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Score	Query Match	Length	DB	ID	Description
1	144	100.0	346	5	BY103030	BY103030 BY103030
2	144	100.0	365	5	BY080676	BY080676 BY080676
3	144	100.0	1123	5	BX376891	BX376891 BX376891
4	144	100.0	1506	9	AY417360	AY417360 Mus sapi
5	144	100.0	1506	9	AY417362	AY417362 Mus muscu
6	144	100.0	3634	3	AK041285	AK041285 Mus muscu
7	144	100.0	3805	3	AK082230	AK082230 Mus muscu
8	144	100.0	3859	3	AK014464	AK014464 Mus muscu
9	144	100.0	3877	3	AK033112	AK033112 Mus muscu

10	144	100.0	3880	3	AK080498	AK080498 Mus muscu
11	144	100.0	4048	3	AK082317	AK082317 Mus muscu
12	144	100.0	4101	3	AK046175	AK046175 Mus muscu
13	143	99.3	4466	3	AK049626	AK049626 Mus muscu
14	138	95.8	461	1	AL700831	AL700831 DKF26861
15	138	95.8	720	4	BG288435	BG288435 602383404
16	137	95.8	813	7	CN224123	CN224123 WLA053E12
17	137	95.8	458	1	AL700814	AL700814 DKF26861
18	130	90.3	611	7	CN484125	CN484125 hw42d08.Y
19	128	88.9	727	6	CA749486	CA749486 UT-M-FY0
20	125	86.8	1001	5	BU128383	BU128383 603133984
21	124	86.1	761	7	CN064831	CN064831 Bg2_p8_p2
22	124	86.1	763	7	CN064511	CN064511 Ag2_p7_p9
23	124	86.1	817	7	CK139305	CK139305 AGENCOURT
24	124	86.1	819	7	CK143759	CK143759 AGENCOURT
25	124	86.1	873	6	CD755522	CD755522 AGENCOURT
26	124	86.1	893	6	CA475966	CA475966 AGENCOURT
27	124	86.1	898	6	CD757678	CD757678 AGENCOURT
28	124	86.1	915	6	CD756050	CD756050 AGENCOURT
29	122	84.7	880	5	BQ733989	BQ733989 AGENCOURT
30	120	83.3	556	1	AL788837	AL788837 AL788837
31	120	83.3	670	6	CA375995	CA375995 654231 NC
32	119	82.6	208	1	AI290317	AI290317 CM020505.X
33	119	82.6	542	2	BE668968	BE668968 159420 MA
34	119	82.6	614	2	AW153854	AW153854 1126d02.Y
35	119	82.6	648	5	BM957312	BM957312 EY76E05.Y
36	119	82.6	705	5	BP434294	BP434294 BP434294
37	119	82.6	712	7	CK681409	CK681409 ZF101-P00
38	119	82.6	751	7	CV480976	CV480976 AGENCOURT
39	119	82.6	765	4	BM006442	BM006442 603615166
40	119	82.6	778	7	CN016970	CN016970 AGENCOURT
41	119	82.6	783	6	CB962018	CB962018 AGENCOURT
42	119	82.6	795	6	CB988213	CB988213 AGENCOURT
43	119	82.6	847	6	CB996886	CB996886 AGENCOURT
44	119	82.6	814	7	CK871931	CK871931 AGENCOURT
45	119	82.6	847	7	CN021417	CN021417 AGENCOURT

#### ALIGNMENTS

RESULT 1  
BY103030  
LOCUS  
DEFINITION  
BY103030  
346 bp mRNA linear EST 07-DEC-2002  
etc. Mus musculus full-length enriched, pooled tissues, adult spleen,  
etc. Mus musculus cDNA clone K630148P20 5', mRNA sequence.

ACCESSION  
BY103030  
VERSION  
BY103030.1 GI:26213647  
KEYWORDS  
SOURCE  
ORGANISM  
Mus musculus (house mouse)

#### REFERENCE

1 (bases 1 to 346)  
Okazaki, Y., Furuno, M., Kasukawa, T., Adachi, J., Bono, H., Kondo, S.,  
Nikaido, I., Osato, N., Saito, R., Suzuki, H., Yamanka, I.,  
Kiyosawa, H., Yagi, K., Tomaru, Y., Hasegawa, Y., Nogami, A.,  
Schombach, C., Gotohori, T., Balderelli, R., Hill, D.P., Bult, C.,  
Hume, D.A., Quackenbush, J., Schriml, L.M., Kanpin, A., Matsuda, H.,  
Batalov, S., Beisel, K.W., Blake, J.A., Brad, D., Brusic, V.,  
Choctha, C., Corbani, L.E., Cousins, S., Dalla, E., Dragan, T.A.,  
Fletcher, C.F., Forrest, A., Frazer, K.S., Gaasterland, T.,  
Gariboldi, M., Gissi, C., Godzik, A., Gough, J., Jarvis, E.D., Kanai, A.,  
Gustincich, S., Hirokawa, N., Jackson, J.J., Jarvis, E.D., Kanai, A.,  
Kawaji, H., Kawasawa, Y., Kedzierski, R.M., King, B.L., Konagaya, A.,  
Kurochkin, I.V., Lee, Y., Lenhard, B., Lyons, P.A., Maglocz, D.R.,  
Maltais, L., Marchionni, L., McKenzie, L., Miki, H., Nagashima, T.,  
Numata, K., Okido, T., Pavan, W.J., Pertea, G., Pesole, G.,  
Petrovsky, N., Pillai, R., Pontius, J.U., Qi, D., Ramchandran, S.,  
Ravasi, T., Reed, J.C., Reed, D.J., Reid, J., Ring, B.Z., Ringwald, M.,  
Santelini, A., Schneider, C., Sempé, C.A., Setou, M., Shimada, K.,  
Sultana, R., Takenaka, Y., Taylor, M.S., Teasdale, R.D., Tomita, M.,  
Verardo, R., Wagner, L., Wahlstedt, C., Wang, Y., Watanabe, Y.,  
Wells, C., Wilming, L.G., Wynshaw-Boris, A., Yanagisawa, M., Yang, I.,

Yang, L., Yuan, Z., Zavolan, M., Zhu, Y., Zimmer, A., Carninci, P., Hayatsu, N., Hirozane-Kishikawa, T., Kono, H., Nakamura, M., Sakazume, N., Sato, K., Shiraki, T., Waki, K., Kawai, J., Aizawa, K., Aikawa, T., Fukuda, S., Hara, A., Hashizume, W., Imotani, K., Ishii, Y., Itoh, M., Kagawa, I., Miyazaki, A., Sakai, K., Sasaki, D., Shibata, K., Shinagawa, A., Yasunishi, A., Yoshino, M., Waterston, R., Lander, E.S., Rogers, J., Birney, E., and Hayashizaki, Y.  
Analysis of the mouse transcriptome based on functional annotation of 60,770 full-length cDNAs  
Nature 420, 563-573 (2002)  
22354683  
12466851  
Contact: Yoshihide Hayashizaki  
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Fax: 81-45-503-9216  
Email: genome-res@gs.c.riken.jp, URL: http://genome.gsc.riken.jp/  
Aizawa, K., Akimura, T., Aikawa, T., Carninci, P., Fukuda, S., Hirozane, T., Imotani, K., Ishii, Y., Itoh, M., Kawai, J., Kono, H., Miyazaki, A., Murata, M., Nakamura, M., Nomura, K., Numazaki, R., Ohno, M., Sakai, K., Sakazume, N., Sasaki, D., Sato, K., Shibata, K., Shiraki, T., Tagami, M., Waki, K., Watabiki, A., Muramatsu, M., and Hayashizaki, Y. Direct Submission  
Computational Analysis of Full-Length Mouse cDNAs Compared with Human Genome Sequences Mamm. Genome. 12, 673-677 (2001)  
Normalization and subtraction of cap-trapper-selected cDNAs to prepare full-length cDNA libraries for rapid discovery of new genes. Genome Res. 10 (10), 1617-1630 (2000)  
RIKEN integrated sequence analysis (RISA) system-384-format sequencing pipeline with 384 multichannel sequencer. Genome Res. 10 (11), 1757-1771 (2000)  
Computer-based methods for the mouse full-length cDNA encyclopedia: real-time sequence clustering for construction of a nonredundant cDNA library. Genome Res. 11 (2), 281-289 (2001)  
cDNA library was prepared and sequenced in Mouse Genome Encyclopedia Project of Genome Exploration Research Group in Riken Division of Experimental Animal Research in Riken contributed to please visit our web site (http://genome.gsc.riken.go.jp) for further details.

## FEATURES

source  
1.346  
Location/Qualifiers  
/organism="Mus musculus"  
/mol\_type="mRNA"  
/db\_xref="taxon:10090"  
/clone\_lib="RIKEN full-length enriched, pooled tissues, adult spleen, etc."  
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ORIGIN  
Alignment Scores:  
Pred. No.: 1.7e-13

Length: 346

Score: 144.00  
Percent Similarity: 100.00%  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
DB: 5  
Gaps: 0

US-10-726-967a-52 (1-28) x BY103030 (1-346)

QY 1 GYTYTYTVAAGLUMETThrValG1SerProProGlnThrLeuAn1LeuVal1asp 20

Db 86 GGCTACTATGTGGAGATATACCGTACGACCCGCCAGAGCTCAACATCTCGTGAC 145

QY 21 ThrG1SerSerAanPheAlaVal 28

Db 146 ACGGCACTAGTAACCTTGACAGT 169

RESULT 2  
LOCUS BY080676 365 bp mRNA linear EST 07-DEC-2002

DERIVATION BY080676 RIKEN full-length enriched, 16 days embryo whole body Mus

ACCESSION BY080676

VERSION BY080676.1 GI:26191219

KEYWORDS EST.

SOURCE Mus musculus (house mouse)

ORGANISM Mus musculus

REFERENCE 1 (bases 1 to 365)

AUTHORS Okazaki, Y., Furuno, M., Kasukawa, T., Aach, J., Bono, H., Kondo, S.,

Nikaido, I., Oosato, N., Saito, R., Suzuki, H., Yamanaka, I.,

Kiyosawa, H., Yagi, K., Tomaru, Y., Hasegawa, Y., Nogami, A.,

Schobach, C., Gojobori, T., Balderelli, L.M., Kanaph, A., Bulc, C.,

Hume, D.A., Quackenbush, J., Schriml, L.M., Kanaph, A., Matsuda, H.,

Batalov, S., Betsel, K.W., Blake, J.A., Brad, D., Brusic, V.,

Chothia, C., Corbani, L.E., Cousins, S., Dalla, E., Dargatz, T.A.,

Fletcher, C.F., Forrest, A., Fraser, K.S., Gaasterland, T.,

Gariboldi, M., Gissi, C., Godzik, A., Gough, J., Grimond, S.,

Guarinch, S., Hirokawa, N., Jackson, I.J., Jerns, E.D., Kanai, A.,

Kawaji, H., Kawasawa, Y., Kedierski, R.M., King, B.L., Konagaya, A.,

Kurochkin, I.V., Lee, Y., Lennard, B., Lyons, P.A., Maglott, D.R.,

Maitals, L., Marchionni, L., McKenzie, L., Miki, H., Nagashima, T.,

Nimata, K., Okido, T., Pavan, W.J., Perce, G., Pesole, G.,

Petrovsky, N., Pillai, R., Pontius, J.U., Qi, D., Ramchandran, S.,

Ravasi, T., Reed, J.C., Reed, D.J., Reid, J., Ring, B.Z., Ringwald, M.,

Sandelin, A., Schneider, C., Semp, C.A., Setu, M., Shimada, K.,

Sultana, R., Takenaka, Y., Taylor, M.S., Teasdale, R.D., Tomita, M.,

Verardo, R., Wagner, L., Wahlestedt, C., Wang, Y., Watanabe, Y.,

Wells, C., Wilting, L.G., Wynshaw-Boris, A., Yanagisawa, M., Yang, I.,

Yang, L., Yuan, Z., Zavolan, M., Zhu, Y., Zimmer, A., Carninci, P.,

Hayatsu, N., Hirozane-Kishikawa, T., Kono, H., Nakamura, M.,

Sakazume, N., Sato, K., Shiraki, T., Waki, K., Kawai, J., Aizawa, K.,

Aikawa, T., Fukuda, S., Hara, A., Hashizume, W., Imotani, K., Ishii, Y.,

Itoh, M., Kagawa, I., Miyazaki, A., Sakai, K., Sasaki, D., Shibata, K.,

Shinagawa, A., Yasunishi, A., Yoshino, M., Waterston, R., Lander, E.S.,

Rogers, J., Birney, E., and Hayashizaki, Y.

Analysis of the mouse transcriptome based on functional annotation

of 60,770 full-length cDNAs

Nature 420, 563-573 (2002)

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12466851

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Fax: 81-45-503-9216  
Email: genome-res@gs.c.riken.jp, URL: http://genome.gsc.riken.jp/  
Aizawa, K., Akimura, T., Aikawa, T., Carninci, P., Fukuda, S., Hirozane, T., Imotani, K., Ishii, Y., Itoh, M., Kawai, J., Kono, H., Miyazaki, A., Murata, M., Nakamura, M., Nomura, K., Numazaki, R., Ohno, M., Sakai, K., Sakazume, N., Sasaki, D., Sato, K., Shibata, K.,



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/gene="BACE"  
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## ORIGIN

## Alignment Scores:

Pred. No.: 1.01e-12 Length: 1506  
Score: 144.00 Matches: 28  
Percent Similarity: 100.00% Conservative: 0  
Best Local Similarity: 100.00% Mismatches: 0  
Query Match: 100.00% Indels: 0  
DB: 9 Gaps: 0

US-10-726-967a-52 (1-28) x AY417360 (1-1506)

Qy 1 G|Y|Y|T|Y|V|A|G|U|E|T|H|V|A|G|Y|S|E|P|P|R|O|G|I|N|T|H|L|E|U|A|E|N|I|L|E|U|V|A|L|A|S|P 20

Db 220 G|G|C|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|T|G|G|C|A|G|C|C|C|C|C|C|C|G|A|G|C|T|C|A|C|A|C|T|C|T|G|G|T|G|A|C 279

Qy 21 T|H|G|Y|S|E|S|E|R|A|N|P|H|E|A|L|A|V|A|L 28

Db 280 A|C|G|G|C|A|G|C|A|G|T|T|G|C|A|G|T|G 303

## RESULT 5

AY417362

LOCUS

AY417362 Mus musculus BACE gene, VIRTUAL TRANSCRIPT, partial sequence.

ACCESSION

AY417362

VERSION

AY417362.1 GI:39773322

KEYWORDS

GSS.

SOURCE

Mus musculus (house mouse)

ORGANISM

Mus musculus

REFERENCE

AUTHORS

TITLE

JOURNAL

PUBMED

AUTHORS

TITLE

JOURNAL

PUBMED

AUTHORS

TITLE

JOURNAL

PUBMED

AUTHORS

TITLE

JOURNAL

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TITLE

JOURNAL

PUBMED

AUTHORS

TITLE

JOURNAL

PUBMED

US-10-726-967a-52 (1-28) x AY417362 (1-1506)

Qy 1 G|Y|Y|T|Y|V|A|G|U|E|T|H|V|A|G|Y|S|E|P|P|R|O|G|I|N|T|H|L|E|U|A|E|N|I|L|E|U|V|A|L|A|S|P 20

Db 220 G|G|C|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|T|G|G|C|A|G|C|C|C|C|C|C|C|G|A|G|C|T|C|A|C|A|C|T|C|T|G|G|T|G|A|C 279

Qy 21 T|H|G|Y|S|E|S|E|R|A|N|P|H|E|A|L|A|V|A|L 28

Db 280 A|C|G|G|C|A|G|C|A|G|T|T|G|C|A|G|T|G 303

## RESULT 5

AK041285

LOCUS

DEFINITION

AK041285

ACCESSION

AK041285

VERSION

AK041285.1 GI:26334342

KEYWORDS

HTC; CAP trapper.

SOURCE

Mus musculus (house mouse)

ORGANISM

Mus musculus

REFERENCE

AUTHORS

TITLE

JOURNAL

PUBMED

AUTHORS

TITLE

JOURNAL

PUBMED

AUTHORS

TITLE

JOURNAL

PUBMED

AUTHORS

TITLE

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TITLE

US-10-726-967a-52 (1-28) x AY417362 (1-1506)

Qy 1 G|Y|Y|T|Y|V|A|G|U|E|T|H|V|A|G|Y|S|E|P|P|R|O|G|I|N|T|H|L|E|U|A|E|N|I|L|E|U|V|A|L|A|S|P 20

Db 220 G|G|C|T|A|C|T|A|G|T|G|A|G|A|T|G|A|C|C|G|T|G|G|C|A|G|C|C|C|C|C|C|C|G|A|G|C|T|C|A|C|A|C|T|C|T|G|G|T|G|A|C 279

Qy 21 T|H|G|Y|S|E|S|E|R|A|N|P|H|E|A|L|A|V|A|L 28

Db 280 A|C|G|G|C|A|G|C|A|G|T|T|G|C|A|G|T|G 303

## RESULT 5

AK041285

LOCUS

DEFINITION

AK041285

ACCESSION

AK041285

VERSION

AK041285.1 GI:26334342

KEYWORDS

HTC; CAP trapper.

SOURCE

Mus musculus (house mouse)

ORGANISM

Mus musculus

REFERENCE

AUTHORS

TITLE

JOURNAL

PUBMED

AUTHORS

TITLE

JOURNAL

PUBMED

AUTHORS

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AUTHORS

TITLE







AAITSEDFKFGNSWMEGILGLAYAEIARPDLSLPPFDLVCQTHIPNIFSLQCGA  
GPILOTETALASVGSMTIGGIDHSIYTSGLSYTPRREMYEIVIRVEIGODLKM  
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## ORIGIN

## Alignment Scores:

Pred. No.:	3,16e-12	Length:	3859
Score:	144.00	Matches:	28
Percent Similarity:	100.00%	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	100.00%	Indels:	0
DB:	3	Gaps:	0

US-10-726-967a-52 (1-28) x AK014464 (1-3859)

Qy 1 GIVTYTYVValGImwethrValGIsrProPcGInThrlenuAnlleuValAap 20

Db 649 GGCCTACTATGTGAGATGACCTGAGCAGCCCCCACAACGCTCAACCTCCTGTCGAC 708

Qy 21 ThrGlySerSerAspPheAlaVal 28

Db 709 ACCGGCAGTACTGTTCTTCAGTGTG 732

RESULT 9 AK033112 3877 bp mRNA linear HTC 03-APR-2004

LOCUS AK033112 Mus musculus 15 days embryo male testis cDNA, RIKEN full-length  
DEFINITION enriched library, clone:8030431G04 product:beta-site APP cleaving  
enzyme, full insert sequence.

ACCESSION AK033112 GI:26328834

VERSION AK033112.1

KEYWORDS HTC; CAP trapper.

SOURCE Mus musculus (house mouse)

ORGANISM Mus musculus

REFERENCE 1 Carninci, P. and Hayashizaki, Y.

AUTHORS High-efficiency full-length cDNA cloning

JOURNAL Meth. Enzymol. 303, 19-44 (1999)

MEDLINE 99279253

PUBMED 10349636

REFERENCE 2 Carninci, P., Shibata, Y., Hayatsu, N., Sugahara, Y., Shibata, K.,

AUTHORS Itoh, M., Kono, H., Okazaki, Y., Muramatsu, M. and Hayashizaki, Y.

JOURNAL Normalization and subtraction of cap-trapper-selected cDNAs to

MEDLINE 20499374

PUBMED 11042159

REFERENCE 3 Shibata, K., Itoh, M., Aizawa, K., Nagaoka, S., Sasaki, N., Carninci, P.,

AUTHORS Kono, H., Akiyama, J., Nishi, K., Katsunari, T., Tashiro, H., Itoh, M.,

JOURNAL Sumi, N., Ishii, Y., Nakamura, S., Hazama, M., Nishine, T., Harada, A.,

MEDLINE Yamamoto, R., Matsumoto, H., Sakaguchi, S., Ikegami, T., Kashiwagi, K.,

PUBMED Fujisake, S., Inoue, K., Togawa, Y., Izawa, M., Ohara, E., Watabiki, M.,

REFERENCE Yonekura, Y., Ishikawa, T., Ozawa, K., Tanaka, T., Matsunaga, S., Kawai, J.,

AUTHORS Okazaki, Y., Muramatsu, M., Inoue, Y., Kira, A. and Hayashizaki, Y.

TITLE RIKEN integrated sequence analysis (RISA) system-384-format

JOURNAL RIKEN integrated sequence analysis (RISA) system-384-format

MEDLINE sequencing pipeline with 384 multicapillary sequencer

PUBMED Genome Res. 10 (11), 1757-1771 (2000)

REFERENCE 4 The RIKEN Genome Exploration Research Group Phase II Team and the

AUTHORS FANTOM Consortium.

TITLE Functional annotation of a full-length mouse cDNA collection

JOURNAL Nature 409, 685-690 (2001)

PUBMED 11076861

REFERENCE 5 The FANTOM Consortium and the RIKEN Genome Exploration Research

AUTHORS

## TITLE Group Phase I &amp; II Team.

JOURNAL Analysis of the mouse transcriptome based on functional annotation

AUTHORS of 60,770 full-length cDNAs

6 (bases 1 to 3877) Nature 420, 563-573 (2002)

Adachi, J., Aizawa, K., Akimura, T., Arakawa, T., Bono, H., Carninci, P.,

Fukuda, S., Furuno, M., Hanagaki, T., Hara, A., Hashizume, M.,

Hayashida, K., Hayatsu, N., Hiramoto, K., Hirooka, T., Hirozane, T.,

Horii, F., Imotani, K., Ishii, Y., Itoh, M., Kagawa, I., Kasukawa, T.,

Kato, H., Kawai, J., Kojima, Y., Kondo, S., Kono, H., Kouda, M.,

Koyama, S., Kurihara, C., Matsuyama, T., Miyazaki, A., Murata, M.,

Nakamura, Y., Nishi, K., Nomura, K., Nunazaki, R., Ohno, M., Ohnaka, N.,

Okazaki, Y., Saito, R., Saitoh, H., Sakai, C., Sakai, K., Sakazume, N.,

Sano, H., Sasaki, D., Shibata, K., Shinagawa, A., Shiraki, T.,

Sogabe, Y., Tagami, M., Tagawa, A., Takahashi, F., Takaku-Akahira, S.,

Takeda, Y., Tanaka, T., Tomaru, A., Toya, T., Yasunishi, A.,

Muramatsu, M. and Hayashizaki, Y.

Direct Submission

Submitted (16-JUL-2001) Yoshihide Hayashizaki, The Institute of

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Exploration Research Group, RIKEN Genomic Sciences Center (GSC),

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Kanagawa 230-0045, Japan (E-mail: genome-resgsc.riken.jp,

URL: http://genome.gsc.riken.jp/, Tel: 81-45-503-9222,

Fax: 81-45-503-9216)

CDNA library was prepared and sequenced in Mouse Genome

Encyclopedia Project of Genome Exploration Research Group in Riken

Genomic Sciences Center and Genome Science Laboratory in RIKEN.

Division of Experimental Animal Research in Riken contributed to

prepare mouse tissues.

Please visit our web site for further details.

URL: http://genome.gsc.riken.jp/

URL: http://fantom.gsc.riken.jp/

Location/Qualifiers

1. 3877

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/mol\_type="mRNA"

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## ORIGIN

## Alignment Scores:

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Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	100.00%	Indels:	0
DB:	3	Gaps:	0

US-10-726-967a-52 (1-28) x AK033112 (1-3877)

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 Db 669 GCCTACTGTCGAGATGACCGTGAAGCCGCCCCACAGCGCTCAACATCTCTGTGGAC 728  
 Qy 21 ThrGlySerSerAsnPhaIaVal 28  
 Db 729 ACGGCGAGTGAATCTTGCAAGTG 752

RESULT 10  
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 Locus AK080498  
 Definition Mus musculus 7 days neonate cerebellum cDNA, RIKEN full-length  
 enriched library, clone:A730059K08 product:beta-site APP cleaving  
 enzyme, full insert sequence.  
 Accession AK080498  
 Version AK080498.1 GI:26099278  
 Keywords HTC; CAP trapper.  
 Source Mus musculus (house mouse)  
 Organism Mus musculus  
 Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.

REFERENCE  
 AUTHORS Carninci, P. and Hayashizaki, Y.  
 TITLE High-efficiency full-length cDNA cloning  
 JOURNAL Meth. Enzymol. 303, 19-44 (1999)  
 MEDLINE 99279253  
 PUBMED 10349636

AUTHORS 2  
 Carninci, P., Shibata, Y., Hayatsu, N., Sugahara, Y., Shibata, K.,  
 Itoh, M., Kono, H., Okazaki, Y., Muramatsu, M. and Hayashizaki, Y.  
 TITLE Normalization and subtraction of cap-trapper-selected cDNAs to  
 JOURNAL Prepare full-length cDNA libraries for rapid discovery of new genes  
 MEDLINE Genome Res. 10 (10), 1617-1630 (2000)  
 PUBMED 20493374  
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3  
 Shibata, K., Itoh, M., Aizawa, K., Nagaoka, S., Sasaki, N., Carninci, P.,  
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 Fujiwara, S., Inoue, K., Togawa, Y., Izawa, M., Ohara, E., Watanabe, M.,  
 Yoneda, Y., Ishikawa, T., Ozawa, K., Tanaka, T., Matsubara, S., Kawai, J.,  
 Okazaki, Y., Muramatsu, M., Inoue, Y., Kita, A. and Hayashizaki, Y.  
 TITLE RIKEN integrated sequence analysis (RISA) system-384-format  
 JOURNAL sequencing pipeline with 384 multicapillary sequencer  
 MEDLINE Genome Res. 10 (11), 1757-1771 (2000)  
 PUBMED 20530913  
 11076861

4  
 The RIKEN Genome Exploration Research Group Phase II Team and the  
 PANTOM Consortium.  
 TITLE Functional annotation of a full-length mouse cDNA collection  
 JOURNAL Nature 409, 685-690 (2001)  
 MEDLINE 11042159

5  
 The FANTOM Consortium and the RIKEN Genome Exploration Research  
 Group Phase I & II Team.  
 TITLE Analysis of the mouse transcriptome based on functional annotation  
 JOURNAL Nature 420, 563-573 (2002)  
 MEDLINE 6 (bases 1 to 3880)  
 PUBMED 120530913

6  
 Adachi, J., Aizawa, K., Akimura, T., Arikawa, T., Bono, H., Carninci, P.,  
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 Takeda, Y., Tanaka, T., Tomaru, A., Toya, T., Yasunishi, A.,  
 Muramatsu, M. and Hayashizaki, Y.

TITLE Direct Submission  
 JOURNAL Submitted (16-Apr-2002) Yoshihide Hayashizaki, The Institute of  
 Physical and Chemical Research (RIKEN), Laboratory for Genome  
 Exploration Research Group, RIKEN Genomic Sciences Center (GSC),  
 Riken Yokohama Institute, 1-7-22 Suehiro-cho, Tsurumi-ku, Yokohama,  
 Kanagawa 230-0045, Japan (E-mail: genome-res@gsc.riken.jp,  
 URL: http://genome.gsc.riken.jp/, Tel: 81-45-503-9222,  
 Fax: 81-45-503-9216)  
 COMMENT cDNA library was prepared and sequenced in Mouse Genome  
 Encyclopedia Project of Genome Exploration Research Group in Riken  
 Genomic Sciences Center and Genome Science Laboratory in Riken.  
 Division of Experimental Animal Research in Riken contributed to  
 prepare mouse libraries.  
 Please visit our web site for further details.  
 URL: http://genome.gsc.riken.jp/  
 URL: http://fantom.gsc.riken.jp/  
 URL: http://location.qualifiers

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ORIGIN  
 Alignment Scores:  
 Pred. No.: 3,18e-12 Length: 3880  
 Score: 144.00 Matches: 28  
 Percent Similarity: 100.00% Conservative: 0  
 Best Local Similarity: 100.00% Mismatches: 0  
 Query Match: 100.00% Indels: 0  
 DB: 3 Gaps: 0  
 US-10-726-967a-52 (1-28) x AK080498 (1-3880)

Qy 1 GlyTyrValGluMetThrValGlySerProProGlnThrLeuAsnIleLeuValAsp 20  
 Db 671 GCCTACTGTCGAGATGACCGTGAAGCCGCCCCACAGCGCTCAACATCTCTGTGGAC 730

Qy 21 ThrGlySerSerAsnPhaIaVal 28  
 Db 731 ACGGCGAGTGAATCTTGCAAGTG 754

RESULT 11  
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 Locus AK082317  
 Definition Mus musculus 0 day neonate cerebellum cDNA, RIKEN full-length  
 enriched library, clone:C230037E16 product:beta-site APP cleaving  
 enzyme, full insert sequence.  
 Accession AK082317  
 Version AK082317.1 GI:26349644  
 Keywords HTC; CAP trapper.  
 Source Mus musculus (house mouse)  
 Organism Mus musculus  
 Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.

REFERENCE  
 AUTHORS Carninci, P. and Hayashizaki, Y.  
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 Itoh, M., Kono, H., Okazaki, Y., Muramatsu, M. and Hayashizaki, Y.



TITLE	Normalization and subtraction of cap-trapper-selected cDNAs to prepare full-length cDNA libraries for rapid discovery of new genes
JOURNAL	Genome Res. 10 (10), 1617-1630 (2000)
MEDLINE	20499374
PUBMED	11042159
REFERENCE	
AUTHORS	3 Shibata, K., Itoh, M., Aizawa, K., Nagaoka, S., Sasaki, N., Carninci, P., Komano, H., Akiyama, J., Nishi, K., Kitesuna, T., Tashiro, H., Itoh, M., Sumi, N., Ishii, Y., Nakamura, S., Hazama, M., Nishine, T., Harada, A., Yamamoto, R., Matsumoto, H., Sakaguchi, S., Ikegami, T., Kashiwagi, K., Fujiwaka, S., Inoue, K., Togawa, Y., Izawa, M., Ohara, E., Watabiki, M., Yoneda, Y., Ishikawa, T., Ozawa, K., Tanaka, T., Matsunura, S., Kawai, J., Okazaki, Y., Muramatsu, M., Inoue, Y., Kira, A. and Hayashizaki, Y.
TITLE	RIKEN integrated sequence analysis (RISA) system-384-format
JOURNAL	Genome Res. 10 (11), 1757-1771 (2000)
MEDLINE	20530913
PUBMED	11076861
REFERENCE	
AUTHORS	4 The RIKEN Genome Exploration Research Group Phase II Team and the PANTOM Consortium.
TITLE	Functional annotation of a full-length mouse cDNA collection
JOURNAL	Nature 409, 685-690 (2001)
MEDLINE	
PUBMED	5 The PANTOM Consortium and the RIKEN Genome Exploration Research Group Phase I & II Team.
REFERENCE	Analysis of the mouse transcriptome based on functional annotation of 60,770 full-length cDNAs
AUTHORS	6 (bases 1 to 4048) Nature 420, 563-573 (2002)
TITLE	Direct Submission
JOURNAL	Submitted (16-APR-2002) Yoshihide Hayashizaki, The Institute of Physical and Chemical Research (RIKEN), Laboratory for Genome Exploration Research Group, RIKEN Genomic Sciences Center (GSC), RIKEN Yokohama Institute; 1-7-22 Suehito-cho, Tsurumi-Ku, Yokohama, Kanagawa 230-0045, Japan (E-mail: genome-res@gsc.riken.jp, URL: http://genome.gsc.riken.jp/, Tel: 81-45-503-9222, Fax: 81-45-503-9216)
MEDLINE	
PUBMED	
REFERENCE	
AUTHORS	2 cDNA library was prepared and sequenced in Mouse Genome Encyclopaedia Project of Genome Exploration Research Group in Riken Genomic Sciences Center and Genome Science Laboratory in RIKEN.
TITLE	Division of Experimental Animal Research in Riken contributed to prepare mouse tissues.
JOURNAL	Please visit our web site for further details.
MEDLINE	URL: http://genome.gsc.riken.jp/
PUBMED	URL: http://fancom.gsc.riken.jp/.
REFERENCE	Location/Qualifiers
AUTHORS	1. 4048 /organism="Mus musculus" /mol_type="mRNA" /strain="C57BL/6J" /db_xref="PANTOM:DB:C230037E16" /db_xref="taxon:10090" /clone="C230037E16" /issue_type="cerebellum" /clone_id="RIKEN full-length enriched mouse cDNA library" /dev_stage="0 day neonate" 451, _1854 /note="unlabeled protein product; beta-site APP cleaving enzyme (MGI:1346542, GBNM_011792, evidence: BLASTN, 98%, match=3874)
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PUBMED	
REFERENCE	
AUTHORS	
TITLE	Alignment Scores:
JOURNAL	Pred. No.: 3,35e-12 Length: 4048
MEDLINE	Score: 144.00 Matches: 28
PUBMED	Percent Similarity: 100.00% Conservative: 0
REFERENCE	Best Local Similarity: 100.00% Mismatches: 0
AUTHORS	Query Match: 100.00% Indels: 0
TITLE	DB: 3 Gaps: 0
JOURNAL	US-10-726-967A-52 (1-28) x AK082317 (1-4048)
MEDLINE	
PUBMED	
REFERENCE	
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MEDLINE	ACCESSION AK046175.1 GI:26337868 VERSION AK046175.1 GI:26337868 KEYWORDS HTC; CAP trapper. SOURCE Mus musculus (house mouse) ORGANISM Mus musculus
PUBMED	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
REFERENCE	1 Carninci, P. and Hayashizaki, Y. High-efficiency full-length cDNA cloning Meth. Enzymol. 303, 19-44 (1999)
AUTHORS	2 Carninci, P., Shibata, Y., Hayatsu, N., Sugahara, Y., Shibata, K., Itoh, M., Komano, H., Okazaki, Y., Muramatsu, M. and Hayashizaki, Y.
TITLE	Normalization and subtraction of cap-trapper-selected cDNAs to prepare full-length cDNA libraries for rapid discovery of new genes
JOURNAL	Genome Res. 10 (10), 1617-1630 (2000)
MEDLINE	20499374
PUBMED	11042159
REFERENCE	
AUTHORS	3 Shibata, K., Itoh, M., Aizawa, K., Nagaoka, S., Sasaki, N., Carninci, P., Komano, H., Akiyama, J., Nishi, K., Kitesuna, T., Tashiro, H., Itoh, M., Sumi, N., Ishii, Y., Nakamura, S., Hazama, M., Nishine, T., Harada, A., Yamamoto, R., Matsumoto, H., Sakaguchi, S., Ikegami, T., Kashiwagi, K., Fujiwaka, S., Inoue, K., Togawa, Y., Izawa, M., Ohara, E., Watabiki, M., Yoneda, Y., Ishikawa, T., Ozawa, K., Tanaka, T., Matsunura, S., Kawai, J., Okazaki, Y., Muramatsu, M., Inoue, Y., Kira, A. and Hayashizaki, Y.
TITLE	RIKEN integrated sequence analysis (RISA) system-384-format
JOURNAL	Genome Res. 10 (11), 1757-1771 (2000)
MEDLINE	20530913
PUBMED	11076861

REFERENCE  
AUTHORS  
4  
The RIKEN Genome Exploration Research Group Phase II Team and the  
PANTOM Consortium.  
TITLE  
Functional annotation of a full-length mouse cDNA collection  
JOURNAL  
Nature 409, 685-690 (2001)  
REFERENCE  
AUTHORS  
5  
The PANTOM Consortium and the RIKEN Genome Exploration Research  
Group Phase I & II Team.  
TITLE  
Analysis of the mouse transcriptome based on functional annotation  
of 60,770 full-length cDNAs  
JOURNAL  
Nature 420, 563-573 (2002)  
REFERENCE  
AUTHORS  
6 (bases 1 to 4101)  
Adachi, J., Aizawa, K., Akimura, T., Arakawa, T., Bono, H., Carninci, P.,  
Fukuda, S., Furuno, M., Hanagaki, T., Hara, A., Hashizume, W.,  
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Takeda, Y., Tanaka, T., Tomaru, A., Toyota, T., Yasunishi, A.,  
Muramatsu, M. and Hayashizaki, Y.  
TITLE  
Direct Submission  
JOURNAL  
Submitted (16-JUL-2001) Yoshihide Hayashizaki, The Institute of  
Physical and Chemical Research (RIKEN), Laboratory for Genome  
Exploration Research Group, RIKEN Genomic Sciences Center (GSC),  
RIKEN Yokohama Institute, 1-7-22 Suehiro-cho, Tsurumi-ku, Yokohama,  
Kanagawa 230-0045, Japan (E-mail: genome-res@gscc.riken.jp,  
URL: http://genome.gsc.riken.jp/, Tel: 81-45-503-9222,  
Fax: 81-45-503-9216)  
COMMENT  
cDNA library was prepared and sequenced in Mouse Genome  
Encyclopedia Project of Genome Exploration Research Group in Riken  
Genomic Sciences Center and Genome Science Laboratory in RIKEN.  
Division of Experimental Animal Research in Riken contributed to  
prepare mouse tissues.  
Please visit our web site for further details.  
URL: http://genome.gsc.riken.jp/  
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Score: 144.00  
Percent Similarity: 100.00%  
Best Local Similarity: 100.00%  
Query Match: 100.00%  
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Gaps: 0  
US-10-726-967a-52 (1-28) x AK046175 (1-4101)  
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DEFINITION  
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VERSION  
KEYWORDS  
SOURCE  
ORGANISM  
Mus musculus (house mouse)  
HTC; CAP trapper.  
AK049626  
Mus musculus  
Eukaryote; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
1  
Carninci, P. and Hayashizaki, Y.  
TITLE  
High-efficiency full-length cDNA cloning  
JOURNAL  
Meth. Enzymol. 303, 19-44 (1999)  
REFERENCE  
AUTHORS  
2  
Carninci, P., Shibata, Y., Hayatsu, N., Sugahara, Y., Shibata, K.,  
Itoh, M., Konno, H., Okazaki, Y., Muramatsu, M. and Hayashizaki, Y.  
Normalization and subtraction of cap-trapper-selected cDNAs to  
prepare full-length cDNA libraries for rapid discovery of new genes  
Genome Res. 10 (10), 1617-1630 (2000)  
JOURNAL  
MEDLINE  
20499374  
PUBMED  
11042159  
REFERENCE  
AUTHORS  
3  
Shibata, K., Itoh, M., Aizawa, K., Nagaoka, S., Sasaki, N., Carninci, P.,  
Konno, H., Akiyama, J., Nishi, K., Kutsuna, T., Tashiro, H., Itoh, M.,  
Sumi, N., Iehi, Y., Nakamura, S., Hazama, M., Nishine, T., Harada, A.,  
Yamamoto, R., Matsumoto, H., Sakaguchi, S., Ikegami, T., Kaehibagi, K.,  
Fujiwaka, S., Inoue, K., Togawa, Y., Izawa, M., Ohara, E., Watabiki, M.,  
Yoneda, Y., Ishikawa, T., Ozawa, K., Tanaka, T., Matsura, S., Kawai, J.,  
Okazaki, Y., Muramatsu, M., Inoue, Y., Kira, A. and Hayashizaki, Y.  
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sequencing pipeline with 384 multicapillary sequencer  
Genome Res. 10 (11), 1757-1771 (2000)  
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MEDLINE  
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The RIKEN Genome Exploration Research Group Phase II Team and the  
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Functional annotation of a full-length mouse cDNA collection  
JOURNAL  
Nature 409, 685-690 (2001)  
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JOURNAL  
Nature 420, 563-573 (2002)  
REFERENCE  
AUTHORS  
6 (bases 1 to 4046)  
Adachi, J., Aizawa, K., Akimura, T., Arakawa, T., Bono, H., Carninci, P.,  
Fukuda, S., Furuno, M., Hanagaki, T., Hara, A., Hashizume, W.,  
Hayashida, K., Hayatsu, N., Hiramoto, K., Hirooka, T., Hirozane, T.,  
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ORIGIN  
Alignment Scores:  
Pred. No.:

3.4e-12 Length: 4101

Koya, S., Kurihara, C., Matsuyama, T., Miyazaki, A., Murata, M., Nakamura, M., Niehi, K., Nomura, K., Numazaki, R., Ohno, M., Ohnaka, N., Okazaki, Y., Saito, R., Saitoh, H., Sakai, C., Sakai, K., Sakazume, N., Sano, H., Sasaki, D., Shibata, K., Shinagawa, A., Shiraki, T., Sogabe, Y., Tagami, M., Tagawa, A., Takahashi, F., Takaku-Kahira, S., Takeda, Y., Tanaka, T., Tomaru, A., Toya, T., Yasunishi, A., Muramatsu, M. and Hayashizaki, Y.

**TITLE**  
Direct Submission

**JOURNAL**  
Submitted (16-JUL-2001) Yoshihide Hayashizaki, The Institute of Physical and Chemical Research (RIKEN), Laboratory for Genome Exploration Research Group, RIKEN Genomic Sciences Center (GSC), RIKEN Yokohama Institute; 1-7-22 Suehiro-cho, Tsurumi-ku, Yokohama, Kanagawa 230-0045, Japan (E-mail: genome-res@gsr.riken.jp, URL: http://genome.gsc.riken.jp/, Tel: 81-45-503-9222, Fax: 81-45-503-9216)

**COMMENT**  
cDNA library was prepared and sequenced in Mouse Genome Encyclopedia Project of Genome Exploration Research Group in Riken Genomic Sciences Center and Genome Science Laboratory in RIKEN. Division of Experimental Animal Research in Riken contributed to prepare mouse tissues.  
Please visit our web site for further details.  
URL: http://genome.gsc.riken.jp/  
URL: http://fantom.gsc.riken.jp/  
Location/Qualifiers

**FEATURES**  
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1. 4046  
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AATTESDKFPIGNSNMGILGLAVAEIARDDELLEPFDSLVRQTHLPNIFSLQCGA  
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DCKENVYDKSIVDSGTNLRLLPKVFAAAYKSIKAASSTKPDGFGFGLQVCMONG  
TTPWNTFPUVSLTLMGEVNTQSPFRITLIPQOYLAPVEDVATSDCKYFAVSOSTGT  
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**ORIGIN**

**Alignment Scores:**

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Score:	143.00	Matches: 27
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Best Local Similarity:	96.43%	Mismatches: 0
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US-10-726-967a-52 (1-28) x AK049626 (1-4046)

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|||||  
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**Qy** 21 ThG1YserSeranphealaval 28  
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RESULT 14  
AL700831

LOCUS  
DEFINITION  
ACCESSION  
VERSION  
KEYWORDS  
SOURCE  
ORGANISM  
REFERENCE  
AUTHORS  
TITLE  
JOURNAL  
COMMENT  
MIPS  
Ingolstaedter Landstr.1, D-85764 Neuberg, Germany  
This is the 5' sequence of the clone insert  
clone from S. Wiemann, Molecular Genome Analysis, German Cancer  
Research Center (DKFZ), Email: s.wiemann@dkfz-heidelberg.de;  
sequenced by GBF (National Research Centre for Biotechnology Ltd.,  
Braunschweig/Germany) within the cDNA sequencing consortium of the  
German Genome Project.  
No 61 sequence available.  
This clone (DKFZp68612411) is available at the RZPD in Berlin.  
Please contact the RZPD: Ressourcenzentrum, Heubnerweg 6, 14059  
Berlin-Charlottenburg, GERMANY; Email: clone@rzpd.de.

**FEATURES**  
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1. 461  
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**ORIGIN**

**Alignment Scores:**

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US-10-726-967a-52 (1-28) x AL700831 (1-461)

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**Qy** 21 ThG1YserSeranphealaval 28  
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Db 130 ACAGGCGAGTAACTTGGCAGTG 153  
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RESULT 15  
BG288435  
LOCUS  
DEFINITION  
ACCESSION  
VERSION  
KEYWORDS  
SOURCE  
ORGANISM  
REFERENCE  
1 (bases 1 to 720)

AUTHORS NIH-MGC <http://mgc.nci.nih.gov/>.  
 TITLE National Institutes of Health, Mammalian Gene Collection (MGC)  
 JOURNAL Unpublished (1999)  
 COMMENT Contact: Robert Strausberg, Ph.D.  
 Email: [cgabbs@mail.nih.gov](mailto:cgabbs@mail.nih.gov)

Tissue Procurement: ATCC  
 cDNA Library Preparation: Life Technologies, Inc.  
 DNA Library Arrayed by: The I.M.A.G.E. Consortium (LIML)  
 DNA Sequencing by: Incyte Genomics, Inc.  
 Clone distribution: MGC clone distribution information can be  
 found through the I.M.A.G.E. Consortium/LIML at:  
<http://image.llnl.gov>  
 Plate: LHAM10367 row: h column: 02  
 High quality sequence start: 8  
 High quality sequence stop: 715.  
 Location/Qualifiers  
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 /lab\_host="DH10B (phage-resistant)"  
 /clone\_idb="NIH MGC 94"  
 /note="Organ: eye; Vector: pCMV-SPORT6; Site\_1: NotI;  
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 Average insert size 3.3 kb. Library enriched for  
 full-length clones and constructed by Life Technologies.  
 Note: this is a NIH\_MGC library."

## FEATURES

source

## ORIGIN

## Alignment Scores:

Pred. No.:	4.05e-12	Length:	720
Score:	138.00	Matches:	27
Percent Similarity:	100.00%	Conservative:	0
Best Local Similarity:	100.00%	Mismatches:	0
Query Match:	95.83%	Indels:	0
DB:	4	Gaps:	0

US-10-726-967a-52 (1-28) x BG288435 (1-720)

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DB	8	TACTATGTGAGATGACCGTAGGCAGCCCCCACAAGACCGCTCAACATCTCTGTGTGACACG	67
QY	22	GlySerSerAspPheAlaVal	28
DB	68	GGCAGTAGTAACTTGCAGTG	88

Search completed: July 27, 2005, 18:17:56  
 Job time : 3160 secs

GenCore version 5.1.6  
Copyright (c) 1993 - 2005 Compugen Ltd.

OM protein - protein search, using sw model

Run on: July 26, 2005, 16:38:55 ; Search time 41 Seconds  
(without alignments)  
50.980 Million cell updates/sec

Title: US-10-726-967A-52

Perfect score: 144

Sequence: 1 GYVEMTVGSPQTILNIVDTGSSNPAV 28

Scoring table:

BLOSUM62  
Gapop 10.0 , Gapext 0.5

Searched: 513545 seqs, 74649064 residues

Total number of hits satisfying chosen parameters: 513545

Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%

Maximum Match 100%  
Listing first 45 summaries

Database :

Issued Patents AA:\*  
1: /cgn2\_6/prodata/1/iaa/5A\_COMB.pep:\*  
2: /cgn2\_6/prodata/1/iaa/5B\_COMB.pep:\*  
3: /cgn2\_6/prodata/1/iaa/6A\_COMB.pep:\*  
4: /cgn2\_6/prodata/1/iaa/6B\_COMB.pep:\*  
5: /cgn2\_6/prodata/1/iaa/6C\_COMB.pep:\*  
6: /cgn2\_6/prodata/1/iaa/backfillseq.pep:\*

Pred. No. is the number of results predicted by chance to have a  
score greater than or equal to the score of the result being printed,  
and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	144	100.0	231	4 US-09-471-669A-107	Sequence 107, App
2	144	100.0	361	4 US-09-724-566A-75	Sequence 75, App1
3	144	100.0	361	4 US-09-471-669A-75	Sequence 75, App1
4	144	100.0	374	4 US-09-724-566A-71	Sequence 71, App1
5	144	100.0	374	4 US-09-471-669A-71	Sequence 71, App1
6	144	100.0	380	4 US-09-471-669A-108	Sequence 108, App
7	144	100.0	390	4 US-09-724-566A-70	Sequence 70, App1
8	144	100.0	390	4 US-09-471-669A-70	Sequence 70, App1
9	144	100.0	395	4 US-09-724-566A-68	Sequence 68, App1
10	144	100.0	395	4 US-09-471-669A-68	Sequence 68, App1
11	144	100.0	401	4 US-09-471-669A-106	Sequence 106, App
12	144	100.0	407	4 US-09-724-566A-58	Sequence 58, App1
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14	144	100.0	419	4 US-09-471-669A-105	Sequence 105, App
15	144	100.0	419	4 US-09-724-566A-57	Sequence 57, App1
16	144	100.0	419	4 US-09-471-669A-57	Sequence 57, App1
17	144	100.0	420	4 US-09-724-566A-60	Sequence 60, App1
18	144	100.0	420	4 US-09-471-669A-60	Sequence 60, App1
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23	144	100.0	425	4 US-09-548-376D-28	Sequence 28, App1
24	144	100.0	425	4 US-09-794-927A-28	Sequence 28, App1
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26	144	100.0	425	4 US-09-795-847B-28	Sequence 28, App1
27	144	100.0	425	4 US-09-869-414-28	Sequence 26, App1

28	144	100.0	425	4 US-09-548-366F-28	Sequence 28, App1
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31	144	100.0	425	4 US-09-806-194A-28	Sequence 28, App1
32	144	100.0	427	4 US-09-471-669A-65	Sequence 65, App1
33	144	100.0	427	4 US-09-548-372D-51	Sequence 51, App1
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39	144	100.0	428	4 US-09-548-373D-51	Sequence 51, App1
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#### ALIGNMENTS

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RESULT 1
US-09-471-669A-107
; Sequence 107, Application US/09471669A
; Patent No. 6810918
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basl, Guribdal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6810918mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McCollogue, Lisa
; APPLICANT: Ekan Pharmaceuticals, Inc.
; TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS
; FILE REFERENCE: 015270-006430US
; CURRENT APPLICATION NUMBER: US/09/471,669A
; CURRENT FILING DATE: 1999-12-24
; PRIOR APPLICATION NUMBER: US 60/114,408
; PRIOR FILING DATE: 1998-12-31
; PRIOR APPLICATION NUMBER: US 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: US 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 108
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 107
; LENGTH: 231
; TYPE: PRT
; ORGANISM: Mus sp.
; OTHER INFORMATION: pbs/Mulimpain E17 Brain #17 construct
US-09-471-669A-107

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Best Local Similarity 100.0%; Pred. No. 6.5e-14;
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RESULT 2
US-09-724-566A-75
; Sequence 75, Application US/09724566A
; Patent No. 6627739
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; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Guripbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6627739mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; TITLE OF INVENTION: Beta-Secretase Enzyme Compositions and
; FILE REFERENCE: 228-US-NEWC2
; CURRENT APPLICATION NUMBER: US/09/724,566A
; PRIOR FILING DATE: 2000-11-28
; PRIOR APPLICATION NUMBER: US 09/501,708
; PRIOR FILING DATE: 2000-02-10
; PRIOR APPLICATION NUMBER: 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: 60/139,172
; PRIOR FILING DATE: 1999-06-15
; SOFTWARE: FastSeq for Windows Version 4.0
; SEQ ID NO 75
; LENGTH: 361
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-724-566A-75

Query Match
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Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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Db 12 GYVEMTVGSPPTLNILVDGSSNFAV 39

RESULT 3
US-09-471-669A-75
; Sequence 75, Application US/09471669A
; Patent No. 6830918
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Guripbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6830918mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; APPLICANT: Elan Pharmaceuticals, Inc.
; TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS
; FILE REFERENCE: 015270-006430US
; CURRENT APPLICATION NUMBER: US/09/471,669A
; PRIOR FILING DATE: 1999-12-24
; PRIOR APPLICATION NUMBER: US 60/114,408
; PRIOR FILING DATE: 1998-12-31
; PRIOR APPLICATION NUMBER: US 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: US 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 108
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 75
; LENGTH: 361
; TYPE: PRT
; ORGANISM: Homo sapiens
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US-09-471-669A-75

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Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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Db 12 GYVEMTVGSPPTLNILVDGSSNFAV 39

RESULT 4
US-09-724-566A-71
; Sequence 71, Application US/09724566A
; Patent No. 6627739
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Guripbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6627739mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; TITLE OF INVENTION: Beta-Secretase Enzyme Compositions and
; FILE REFERENCE: 228-US-NEWC2
; CURRENT APPLICATION NUMBER: US/09/724,566A
; PRIOR FILING DATE: 2000-11-28
; PRIOR APPLICATION NUMBER: US 09/501,708
; PRIOR FILING DATE: 2000-02-10
; PRIOR APPLICATION NUMBER: 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 104
; SOFTWARE: FastSeq for Windows Version 4.0
; SEQ ID NO 71
; LENGTH: 374
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-724-566A-71

Query Match
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Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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Db 29 GYVEMTVGSPPTLNILVDGSSNFAV 56

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; Sequence 71, Application US/09471669A
; Patent No. 6830918
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Guripbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6830918mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; APPLICANT: Elan Pharmaceuticals, Inc.
; TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS
```

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FILE REFERENCE: 015270-006430US
; CURRENT APPLICATION NUMBER: US/09/471,669A
; CURRENT FILING DATE: 1999-12-24
; PRIOR APPLICATION NUMBER: US 60/114,408
; PRIOR FILING DATE: 1998-12-31
; PRIOR APPLICATION NUMBER: US 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: US 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 108
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 71
; LENGTH: 374
; TYPE: PRT
; ORGANISM: Homo sapiens
; US-09-471-669A-71

Query Match          100.0%; Score 144; DB 4; Length 374;
Best Local Similarity 100.0%; Pred. No. 1,1e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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Db 29 GYVENTVGSPPQTNIILVDTGSSNFAV 56

RESULT 6
US-09-471-669A-108
; Sequence 108, Application US/09471669A
; Patent No. 6830918
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6830918mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; APPLICANT: Elan Pharmaceuticals, Inc.
; TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS
; FILE REFERENCE: 015270-006430US
; CURRENT APPLICATION NUMBER: US/09/471,669A
; CURRENT FILING DATE: 1999-12-24
; PRIOR APPLICATION NUMBER: US 60/114,408
; PRIOR FILING DATE: 1998-12-31
; PRIOR APPLICATION NUMBER: US 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: US 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 108
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 108
; LENGTH: 380
; TYPE: PRT
; ORGANISM: Mus sp.
; FEATURE:
; OTHER INFORMATION: PBS/Multipain E17 Brain#15 construct
; US-09-471-669A-108

Query Match          100.0%; Score 144; DB 4; Length 380;
Best Local Similarity 100.0%; Pred. No. 1.2e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 1 GYVENTVGSPPQTNIILVDTGSSNFAV 28
Db 15 GYVENTVGSPPQTNIILVDTGSSNFAV 42

RESULT 7
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US-09-724-566A-70
; Sequence 70, Application US/09724566A
; Patent No. 6627739
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6627739mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; TITLE OF INVENTION: Beta-Secretase Enzyme Compositions and
; FILE REFERENCE: 228-US-NEWC2
; CURRENT APPLICATION NUMBER: US/09/724,566A
; CURRENT FILING DATE: 2000-11-28
; PRIOR APPLICATION NUMBER: US 09/501,708
; PRIOR FILING DATE: 2000-02-10
; PRIOR APPLICATION NUMBER: 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 104
; SOFTWARE: FastSeq for Windows Version 4.0
; SEQ ID NO 70
; LENGTH: 390
; TYPE: PRT
; ORGANISM: Homo sapiens
; US-09-724-566A-70

Query Match          100.0%; Score 144; DB 4; Length 390;
Best Local Similarity 100.0%; Pred. No. 1.2e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 1 GYVENTVGSPPQTNIILVDTGSSNFAV 28
Db 12 GYVENTVGSPPQTNIILVDTGSSNFAV 39

RESULT 8
US-09-471-669A-70
; Sequence 70, Application US/09471669A
; Patent No. 6830918
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6830918mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; APPLICANT: Elan Pharmaceuticals, Inc.
; TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS
; FILE REFERENCE: 015270-006430US
; CURRENT APPLICATION NUMBER: US/09/471,669A
; CURRENT FILING DATE: 1999-12-24
; PRIOR APPLICATION NUMBER: US 60/114,408
; PRIOR FILING DATE: 1998-12-31
; PRIOR APPLICATION NUMBER: US 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: US 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 108
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 70
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LENGTH: 390  
TYPE: PRT  
ORGANISM: Homo sapiens  
US-09-471-669A-70

Query Match  
Best Local Similarity 100.0%; Score 144; DB 4; Length 390;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTLITLIVDTGSSNFAY 28  
DB 12 GYVEMTVGSPPTLITLIVDTGSSNFAY 39

RESULT 9  
US-09-724-566A-68

Sequence 68, Application US/09724566A  
Patent No. 6627739

GENERAL INFORMATION:

APPLICANT: Anderson, John P.

APPLICANT: Basl, Gurigbal

APPLICANT: Doane, Minh Tam

APPLICANT: Frigson, No. 6627739mand

APPLICANT: John, Varghese

APPLICANT: Power, Michael

APPLICANT: Sinha, Sukanto

APPLICANT: Tatsuno, Gwen

APPLICANT: Tung, Jay

APPLICANT: Wang, Shuwen

APPLICANT: McConlogue, Lisa

TITLE OF INVENTION: Beta-Secretase Enzyme Compositions and

FILE REFERENCE: 228-US-NEMC2

CURRENT APPLICATION NUMBER: US/09/724,566A

PRIOR FILING DATE: 2000-11-28

PRIOR APPLICATION NUMBER: US 09/501,708

PRIOR FILING DATE: 2000-02-10

PRIOR APPLICATION NUMBER: 60/119,571

PRIOR FILING DATE: 1999-02-10

PRIOR APPLICATION NUMBER: 60/139,172

PRIOR FILING DATE: 1999-06-15

NUMBER OF SEQ ID NOS: 104

SOFTWARE: PaeSeq for Windows Version 4.0

SEQ ID NO 68

LENGTH: 395

TYPE: PRT

ORGANISM: Homo sapiens

US-09-724-566A-68

Query Match  
Best Local Similarity 100.0%; Score 144; DB 4; Length 395;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTLITLIVDTGSSNFAY 28  
DB 17 GYVEMTVGSPPTLITLIVDTGSSNFAY 44

RESULT 10

US-09-471-669A-68

Sequence 68, Application US/09471669A

Patent No. 6830918

GENERAL INFORMATION:

APPLICANT: Anderson, John P.

APPLICANT: Basl, Gurigbal

APPLICANT: Doane, Minh Tam

APPLICANT: Frigson, No. 6830918mand

APPLICANT: John, Varghese

APPLICANT: Power, Michael

APPLICANT: Sinha, Sukanto

APPLICANT: Tatsuno, Gwen

APPLICANT: Tung, Jay

APPLICANT: Wang, Shuwen

APPLICANT: McConlogue, Lisa  
APPLICANT: Eian Pharmaceuticals, Inc.  
TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS

FILE REFERENCE: 015270-006430US

CURRENT APPLICATION NUMBER: US/09/471,669A

PRIOR FILING DATE: 1999-12-24

PRIOR APPLICATION NUMBER: US 60/114,408

PRIOR FILING DATE: 1998-12-31

PRIOR APPLICATION NUMBER: US 60/119,571

PRIOR FILING DATE: 1999-02-10

PRIOR APPLICATION NUMBER: US 60/139,172

PRIOR FILING DATE: 1999-06-15

NUMBER OF SEQ ID NOS: 108

SOFTWARE: PatentIn Ver. 2.1

SEQ ID NO 68

LENGTH: 395

TYPE: PRT

ORGANISM: Homo sapiens

US-09-471-669A-68

Query Match  
Best Local Similarity 100.0%; Score 144; DB 4; Length 395;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTLITLIVDTGSSNFAY 28  
DB 17 GYVEMTVGSPPTLITLIVDTGSSNFAY 44

RESULT 11

US-09-471-669A-106

Sequence 106, Application US/09471669A

Patent No. 6830918

GENERAL INFORMATION:

APPLICANT: Anderson, John P.

APPLICANT: Basl, Gurigbal

APPLICANT: Doane, Minh Tam

APPLICANT: Frigson, No. 6830918mand

APPLICANT: John, Varghese

APPLICANT: Power, Michael

APPLICANT: Sinha, Sukanto

APPLICANT: Tatsuno, Gwen

APPLICANT: Tung, Jay

APPLICANT: Wang, Shuwen

APPLICANT: McConlogue, Lisa

TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS

FILE REFERENCE: 015270-006430US

CURRENT APPLICATION NUMBER: US/09/471,669A

PRIOR FILING DATE: 1999-12-24

PRIOR APPLICATION NUMBER: US 60/114,408

PRIOR FILING DATE: 1998-12-31

PRIOR APPLICATION NUMBER: US 60/119,571

PRIOR FILING DATE: 1999-02-10

PRIOR APPLICATION NUMBER: US 60/139,172

PRIOR FILING DATE: 1999-06-15

NUMBER OF SEQ ID NOS: 108

SOFTWARE: PatentIn Ver. 2.1

SEQ ID NO 106

LENGTH: 401

TYPE: PRT

ORGANISM: Mus sp.

FEATURE:

OTHER INFORMATION: pbs/MuImPain E17 #14 construct

US-09-471-669A-106

Query Match  
Best Local Similarity 100.0%; Score 144; DB 4; Length 401;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTLITLIVDTGSSNFAY 28  
DB 25 GYVEMTVGSPPTLITLIVDTGSSNFAY 52



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RESULT 12
US-09-724-566A-58
; Sequence 58, Application US/09724566A
; Patent No. 6627739
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6627739mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; TITLE OF INVENTION: Beta-Secretase Enzyme Compositions and
; TITLE OF INVENTION: Methods
; FILE REFERENCE: 228-US-NEWC2
; CURRENT APPLICATION NUMBER: US/09/724,566A
; PRIOR FILING DATE: 2000-11-28
; PRIOR APPLICATION NUMBER: US 09/501,708
; PRIOR FILING DATE: 2000-02-10
; PRIOR APPLICATION NUMBER: 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 104
; SOFTWARE: FastSeq for Windows Version 4.0
; SEQ ID NO 58
; LENGTH: 407
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-724-566A-58

Query Match          100.0%; Score 144; DB 4; Length 407;
Best Local Similarity 100.0%; Pred. No. 1.3e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 GYVENTVGSPPQTNIILVDTGSSNPAV 28
DB      29 GYVENTVGSPPQTNIILVDTGSSNPAV 56

RESULT 13
US-09-471-669A-58
; Sequence 58, Application US/09471669A
; Patent No. 6830918
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6830918mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; APPLICANT: Eilan Pharmaceuticals, Inc.
; TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS
; FILE REFERENCE: 015270-006430US
; CURRENT APPLICATION NUMBER: US/09/471,669A
; PRIOR FILING DATE: 1999-12-24
; PRIOR APPLICATION NUMBER: US 60/114,408
; PRIOR FILING DATE: 1998-12-31
; PRIOR APPLICATION NUMBER: 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: US 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 105
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 105
; LENGTH: 408
; TYPE: PRT
; ORGANISM: Mus sp.
; FEATURE:
; OTHER INFORMATION: PBS/Multipain E17 #11 construct
US-09-471-669A-105

Query Match          100.0%; Score 144; DB 4; Length 408;
Best Local Similarity 100.0%; Pred. No. 1.3e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 GYVENTVGSPPQTNIILVDTGSSNPAV 28
DB      27 GYVENTVGSPPQTNIILVDTGSSNPAV 54

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; NUMBER OF SEQ ID NOS: 108
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 58
; LENGTH: 407
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-471-669A-58

Query Match          100.0%; Score 144; DB 4; Length 407;
Best Local Similarity 100.0%; Pred. No. 1.3e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 GYVENTVGSPPQTNIILVDTGSSNPAV 28
DB      29 GYVENTVGSPPQTNIILVDTGSSNPAV 56

RESULT 14
US-09-471-669A-105
; Sequence 105, Application US/09471669A
; Patent No. 6830918
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6830918mand
; APPLICANT: John, Varghese
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tatsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; APPLICANT: Eilan Pharmaceuticals, Inc.
; TITLE OF INVENTION: BETA-SECRETASE ENZYME COMPOSITIONS AND METHODS
; FILE REFERENCE: 015270-006430US
; CURRENT APPLICATION NUMBER: US/09/471,669A
; PRIOR FILING DATE: 1999-12-24
; PRIOR APPLICATION NUMBER: US 60/114,408
; PRIOR FILING DATE: 1998-12-31
; PRIOR APPLICATION NUMBER: 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: US 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 108
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 105
; LENGTH: 408
; TYPE: PRT
; ORGANISM: Mus sp.
; FEATURE:
; OTHER INFORMATION: PBS/Multipain E17 #11 construct
US-09-471-669A-105

Query Match          100.0%; Score 144; DB 4; Length 408;
Best Local Similarity 100.0%; Pred. No. 1.3e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY      1 GYVENTVGSPPQTNIILVDTGSSNPAV 28
DB      27 GYVENTVGSPPQTNIILVDTGSSNPAV 54

RESULT 15
US-09-724-566A-57
; Sequence 57, Application US/09724566A
; Patent No. 6627739
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P.
; APPLICANT: Basi, Gurigbal
; APPLICANT: Doane, Minh Tam
; APPLICANT: Frigon, No. 6627739mand
; APPLICANT: John, Varghese

```

```
; APPLICANT: Power, Michael
; APPLICANT: Sinha, Sukanto
; APPLICANT: Tarsuno, Gwen
; APPLICANT: Tung, Jay
; APPLICANT: Wang, Shuwen
; APPLICANT: McConlogue, Lisa
; TITLE OF INVENTION: Beta-Secretase Enzyme Compositions and
; TITLE OF INVENTION: Methods
; FILE REFERENCE: 228-US-NEMC2
; CURRENT APPLICATION NUMBER: US/09/724,566A
; CURRENT FILING DATE: 2000-11-28
; PRIOR APPLICATION NUMBER: US 09/501,708
; PRIOR FILING DATE: 2000-02-10
; PRIOR APPLICATION NUMBER: 60/119,571
; PRIOR FILING DATE: 1999-02-10
; PRIOR APPLICATION NUMBER: 60/139,172
; PRIOR FILING DATE: 1999-06-15
; NUMBER OF SEQ ID NOS: 104
; SOFTWARE: FastSeq for Windows Version 4.0
; SEQ ID NO 57
; LENGTH: 419
; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-724-566A-57
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Query Match          100.0%; Score 144; DB 4; Length 419;
Best Local Similarity 100.0%; Pred. No. 1.3e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
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      |||||
Db      74 GYVEMTVGSPPTLNILVDTGSSNFAV 101
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GenCore version 5.1.6  
Copyright (c) 1993 - 2005 Compugen Ltd

OM protein - protein search, using sw model

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Title: US-10-726-967A-52  
Perfect score: 144  
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Published Applications AA:\*

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- 2: /cgn2\_6/ptodataa/2/pubpaa/PTCT\_NEW\_PUB.pdp.\*
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- 9: /cgn2\_6/ptodataa/2/pubpaa/US09A\_PUBCOMB.pdp.\*
- 10: /cgn2\_6/ptodataa/2/pubpaa/US09B\_PUBCOMB.pdp.\*
- 11: /cgn2\_6/ptodataa/2/pubpaa/US09C\_PUBCOMB.pdp.\*
- 12: /cgn2\_6/ptodataa/2/pubpaa/US09\_NEW\_PUB.pdp.\*
- 13: /cgn2\_6/ptodataa/2/pubpaa/US10A\_PUBCOMB.pdp.\*
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- 17: /cgn2\_6/ptodataa/2/pubpaa/US10E\_PUBCOMB.pdp.\*
- 18: /cgn2\_6/ptodataa/2/pubpaa/US10\_NEW\_PUB.pdp.\*
- 19: /cgn2\_6/ptodataa/2/pubpaa/US11A\_PUBCOMB.pdp.\*
- 20: /cgn2\_6/ptodataa/2/pubpaa/US11\_NEW\_PUB.pdp.\*
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- 22: /cgn2\_6/ptodataa/2/pubpaa/US60\_PUBCOMB.pdp.\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	DB	ID	Description
1	144	100.0	28	17	US-10-726-967A-52	Sequence 52, Appl
2	144	100.0	358	17	US-10-872-198-12	Sequence 12, Appl
3	144	100.0	358	17	US-10-872-197A-12	Sequence 12, Appl
4	144	100.0	331	15	US-10-372-473-4	Sequence 4, Appl
5	144	100.0	403	15	US-10-400-273-4	Sequence 4, Appl
6	144	100.0	406	16	US-10-837-021A-2	Sequence 2, Appl
7	144	100.0	406	16	US-10-837-021A-3	Sequence 3, Appl
8	144	100.0	406	16	US-10-837-021A-4	Sequence 4, Appl
9	144	100.0	406	16	US-10-837-021A-5	Sequence 5, Appl
10	144	100.0	408	15	US-10-400-273-5	Sequence 5, Appl
11	144	100.0	411	15	US-10-400-273-1	Sequence 1, Appl

12	144	100.0	411	15	US-10-627-473-19	Sequence 19, Appl
13	144	100.0	411	15	US-10-627-473-19	Sequence 20, Appl
14	144	100.0	414	16	US-10-281-092-9	Sequence 9, Appl
15	144	100.0	417	15	US-10-627-473-21	Sequence 21, Appl
16	144	100.0	425	9	US-09-794-927-28	Sequence 28, Appl
17	144	100.0	425	9	US-09-795-847-28	Sequence 28, Appl
18	144	100.0	425	9	US-09-794-743-28	Sequence 28, Appl
19	144	100.0	425	9	US-09-794-748-28	Sequence 28, Appl
20	144	100.0	425	9	US-09-794-925-28	Sequence 28, Appl
21	144	100.0	425	9	US-09-681-442-28	Sequence 28, Appl
22	144	100.0	425	10	US-09-869-414-28	Sequence 28, Appl
23	144	100.0	425	10	US-10-372-473-3	Sequence 3, Appl
24	144	100.0	425	15	US-10-652-830-28	Sequence 28, Appl
25	144	100.0	425	15	US-10-652-830-28	Sequence 28, Appl
26	144	100.0	425	16	US-10-652-935-28	Sequence 28, Appl
27	144	100.0	425	16	US-10-652-935-28	Sequence 28, Appl
28	144	100.0	425	17	US-10-940-867-28	Sequence 28, Appl
29	144	100.0	425	17	US-10-477-076-28	Sequence 28, Appl
30	144	100.0	428	9	US-09-794-897-51	Sequence 51, Appl
31	144	100.0	428	9	US-09-795-867-51	Sequence 51, Appl
32	144	100.0	428	9	US-09-794-743-51	Sequence 51, Appl
33	144	100.0	428	9	US-09-794-748-51	Sequence 51, Appl
34	144	100.0	428	9	US-09-794-925-51	Sequence 51, Appl
35	144	100.0	428	9	US-09-681-442-51	Sequence 51, Appl
36	144	100.0	428	10	US-09-869-414-51	Sequence 51, Appl
37	144	100.0	428	10	US-09-548-366-51	Sequence 51, Appl
38	144	100.0	428	15	US-10-652-827-51	Sequence 51, Appl
39	144	100.0	428	15	US-10-652-830-51	Sequence 51, Appl
40	144	100.0	428	16	US-10-652-935-51	Sequence 51, Appl
41	144	100.0	428	16	US-10-476-935-51	Sequence 51, Appl
42	144	100.0	428	17	US-10-477-076-51	Sequence 51, Appl
43	144	100.0	432	15	US-10-372-473-2	Sequence 26, Appl
44	144	100.0	432	15	US-09-794-927-26	Sequence 26, Appl
45	144	100.0	433	9	US-09-794-927-26	Sequence 26, Appl

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RESULT 1
US-10-726-967A-52
; Sequence 52, Application US/10726967A
; Publication No. US20050074456A1
; GENERAL INFORMATION:
; APPLICANT: Ballinger, Marcus
; TITLE OF INVENTION: Constructs for Homogenously Processed Preparations of Beta Site
; TITLE OF INVENTION: App-leaving Enzyme
; FILE REFERENCE: 2004345-0021
; CURRENT FILING DATE: 2003-12-02
; NUMBER OF SEQ ID NOS: 110
; SOFTWARE: PatentIn version 3.2
; SEQ ID NO 52
; LENGTH: 28
; TYPE: PRT
; ORGANISM: Artificial
; FEATURE:
; OTHER INFORMATION: Residues 74-101 of human BACE1 preprosequence
US-10-726-967A-52

Query Match          100.0%; Score 144; DB 17; Length 28;
Best Local Similarity 100.0%; Pred. No. 1.8e-14;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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        |||||
Db      1 GYYVMTVGSPPTQILIVDTGSSNPAV 28

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; GENERAL INFORMATION:
; APPLICANT: Ulrich HAUPTS
; APPLICANT: Andre KOLTERMANN
; APPLICANT: Andreas SCHEIDIG
; APPLICANT: Christian VOTSMERER
; APPLICANT: Ulrich KETTLING
; TITLE OF INVENTION: NEW BIOLOGICAL ENTITIES AND USE THEREOF
; FILE REFERENCE: 04156.0002U4
; CURRENT APPLICATION NUMBER: US/10/872,198
; CURRENT FILING DATE: 2004-06-18
; PRIOR APPLICATION NUMBER: 60/543,518
; PRIOR FILING DATE: 2004-02-11
; PRIOR APPLICATION NUMBER: 60/524,960
; PRIOR FILING DATE: 2003-11-25
; PRIOR APPLICATION NUMBER: EP 04003058
; PRIOR FILING DATE: 2004-02-11
; PRIOR APPLICATION NUMBER: EP 03025871
; PRIOR FILING DATE: 2003-11-11
; PRIOR APPLICATION NUMBER: EP 03025851
; PRIOR FILING DATE: 2003-11-10
; PRIOR APPLICATION NUMBER: EP 03013819
; PRIOR FILING DATE: 2003-06-18
; NUMBER OF SEQ ID NOS: 149
; SOFTWARE: PatentIn version 3.1
; SEQ ID NO 12
; LENGTH: 358
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-872-198-12

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Query Match          100.0%; Score 144; DB 17; Length 358;
Best Local Similarity 100.0%; Pred. No. 3.5e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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QY 1 GYVEMTVGSPPTNLIVDTGSSNFAV 28
    |||||
DB 13 GYVEMTVGSPPTNLIVDTGSSNFAV 40

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RESULT 3
US-10-872-197A-12
; Sequence 12, Application US/10872197A
; Publication No. US20050059126A1
; GENERAL INFORMATION:
; APPLICANT: Ulrich HAUPTS
; APPLICANT: Andre KOLTERMANN
; APPLICANT: Andreas SCHEIDIG
; APPLICANT: Christian VOTSMERER
; APPLICANT: Ulrich KETTLING
; TITLE OF INVENTION: NEW BIOLOGICAL ENTITIES AND USE THEREOF
; FILE REFERENCE: 04156.0002U3
; CURRENT APPLICATION NUMBER: US/10/872,197A
; CURRENT FILING DATE: 2004-06-18
; PRIOR APPLICATION NUMBER: 60/524,960
; PRIOR FILING DATE: 2003-11-25
; PRIOR APPLICATION NUMBER: EP 03025871
; PRIOR FILING DATE: 2003-11-11
; PRIOR APPLICATION NUMBER: EP 03013819
; PRIOR FILING DATE: 2003-11-10
; NUMBER OF SEQ ID NOS: 96
; SOFTWARE: PatentIn version 3.1
; SEQ ID NO 12
; LENGTH: 358
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-872-197A-12

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Query Match          100.0%; Score 144; DB 17; Length 358;
Best Local Similarity 100.0%; Pred. No. 3.5e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
QY 1 GYVEMTVGSPPTNLIVDTGSSNFAV 28
    |||||

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```

DB 13 GYVEMTVGSPPTNLIVDTGSSNFAV 40

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RESULT 4
US-10-372-473-4
; Sequence 4, Application US/10372473
; Publication No. US20040005691A1
; GENERAL INFORMATION:
; APPLICANT: Chou, Kuo-Chen
; APPLICANT: Howe, W. Jeffery
; TITLE OF INVENTION: Modified BACE
; FILE REFERENCE: MBHB-01-1766-A
; CURRENT APPLICATION NUMBER: US/10/372,473
; CURRENT FILING DATE: 2003-02-21
; NUMBER OF SEQ ID NOS: 24
; SOFTWARE: PatentIn version 3.2
; SEQ ID NO 4
; LENGTH: 391
; TYPE: PRT
; ORGANISM: Homo sapiens
; FEATURE:
; NAME/KEY: MISC FEATURE
; OTHER INFORMATION: Human beta-secretase.
US-10-372-473-4

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Query Match          100.0%; Score 144; DB 15; Length 391;
Best Local Similarity 100.0%; Pred. No. 3.9e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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QY 1 GYVEMTVGSPPTNLIVDTGSSNFAV 28
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DB 19 GYVEMTVGSPPTNLIVDTGSSNFAV 46

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RESULT 5
US-10-400-273-4
; Sequence 4, Application US/10400273
; Publication No. US20040014194A1
; GENERAL INFORMATION:
; APPLICANT: Beyer, Brian
; APPLICANT: Hammond, Gerald S
; APPLICANT: Reichert, Paul
; APPLICANT: Strickland, Corey
; APPLICANT: Wang, Wenyan
; APPLICANT: Weber, Patricia C
; APPLICANT: Wong, Gwendolyn
; APPLICANT: Zhang, Lili
; TITLE OF INVENTION: BETA-SECRETASE CRYSTALS AND METHODS FOR PREPARING AND USING THE S.
; FILE REFERENCE: JBO1531-K-US
; CURRENT APPLICATION NUMBER: US/10/400,273
; CURRENT FILING DATE: 2003-03-26
; PRIOR APPLICATION NUMBER: 60/367,937
; PRIOR FILING DATE: 2002-03-27
; NUMBER OF SEQ ID NOS: 5
; SOFTWARE: PatentIn version 3.1
; SEQ ID NO 4
; LENGTH: 403
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-400-273-4

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```

Query Match          100.0%; Score 144; DB 15; Length 403;
Best Local Similarity 100.0%; Pred. No. 4e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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```

QY 1 GYVEMTVGSPPTNLIVDTGSSNFAV 28
    |||||
DB 27 GYVEMTVGSPPTNLIVDTGSSNFAV 54

```

```

RESULT 6
US-10-837-021A-2
; Sequence 2, Application US/10837021A

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; Publication No. US20040265965A1
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P
; APPLICANT: McConlogue, Lisa
; APPLICANT: Basl, Guribajal
; APPLICANT: Sinha, Sukarno
; TITLE OF INVENTION: Glycosylation Variants of BACE
; FILE REFERENCE: MBHB-03-268-A
; CURRENT APPLICATION NUMBER: US/10/837,021A
; PRIOR FILING DATE: 2004-04-30
; PRIOR APPLICATION NUMBER: 60/467,509
; NUMBER OF SEQ ID NOS: 12
; SOFTWARE: PatentIn version 3.3
; SEQ ID NO 2
; LENGTH: 406
; TYPE: PRT
; ORGANISM: Artificial
; FEATURE:
; OTHER INFORMATION: Human BACE with asparagine to alanine (N223A) mutation.
US-10-837-021A-2

Query Match          100.0%; Score 144; DB 16; Length 406;
Best Local Similarity 100.0%; Pred. No. 4e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTLNILVDTGSSNFAV 28
DB 29 GYVEMTVGSPPTLNILVDTGSSNFAV 56

RESULT 7
US-10-837-021A-3
; Sequence 3, Application US/10837021A
; Publication No. US20040265965A1
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P
; APPLICANT: McConlogue, Lisa
; APPLICANT: Basl, Guribajal
; APPLICANT: Sinha, Sukarno
; TITLE OF INVENTION: Glycosylation Variants of BACE
; FILE REFERENCE: MBHB-03-268-A
; CURRENT APPLICATION NUMBER: US/10/837,021A
; PRIOR FILING DATE: 2004-04-30
; PRIOR APPLICATION NUMBER: 60/467,509
; NUMBER OF SEQ ID NOS: 12
; SOFTWARE: PatentIn version 3.3
; SEQ ID NO 3
; LENGTH: 406
; TYPE: PRT
; ORGANISM: Artificial
; FEATURE:
; OTHER INFORMATION: Human BACE with serine to isoleucine (S174I) and asparagine to
US-10-837-021A-3

Query Match          100.0%; Score 144; DB 16; Length 406;
Best Local Similarity 100.0%; Pred. No. 4e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTLNILVDTGSSNFAV 28
DB 29 GYVEMTVGSPPTLNILVDTGSSNFAV 56

RESULT 8
US-10-837-021A-4
; Sequence 4, Application US/10837021A
; Publication No. US20040265965A1
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P
; APPLICANT: McConlogue, Lisa
```

```
; APPLICANT: Basl, Guribajal
; APPLICANT: Sinha, Sukarno
; TITLE OF INVENTION: Glycosylation Variants of BACE
; FILE REFERENCE: MBHB-03-268-A
; CURRENT APPLICATION NUMBER: US/10/837,021A
; PRIOR FILING DATE: 2004-04-30
; PRIOR APPLICATION NUMBER: 60/467,509
; NUMBER OF SEQ ID NOS: 12
; SOFTWARE: PatentIn version 3.3
; SEQ ID NO 4
; LENGTH: 406
; TYPE: PRT
; ORGANISM: Artificial
; FEATURE:
; OTHER INFORMATION: Human BACE with serine to isoleucine (S174I), asparagine to
US-10-837-021A-4

Query Match          100.0%; Score 144; DB 16; Length 406;
Best Local Similarity 100.0%; Pred. No. 4e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTLNILVDTGSSNFAV 28
DB 29 GYVEMTVGSPPTLNILVDTGSSNFAV 56

RESULT 9
US-10-837-021A-5
; Sequence 5, Application US/10837021A
; Publication No. US20040265965A1
; GENERAL INFORMATION:
; APPLICANT: Anderson, John P
; APPLICANT: McConlogue, Lisa
; APPLICANT: Basl, Guribajal
; APPLICANT: Sinha, Sukarno
; TITLE OF INVENTION: Glycosylation Variants of BACE
; FILE REFERENCE: MBHB-03-268-A
; CURRENT APPLICATION NUMBER: US/10/837,021A
; PRIOR FILING DATE: 2004-04-30
; PRIOR APPLICATION NUMBER: 60/467,509
; NUMBER OF SEQ ID NOS: 12
; SOFTWARE: PatentIn version 3.3
; SEQ ID NO 5
; LENGTH: 406
; TYPE: PRT
; ORGANISM: Artificial
; FEATURE:
; OTHER INFORMATION: Human BACE with serine to isoleucine (S174I), asparagine to
US-10-837-021A-5

Query Match          100.0%; Score 144; DB 16; Length 406;
Best Local Similarity 100.0%; Pred. No. 4e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTLNILVDTGSSNFAV 28
DB 29 GYVEMTVGSPPTLNILVDTGSSNFAV 56

RESULT 10
US-10-400-273-5
; Sequence 5, Application US/10400273
; Publication No. US20040014194A1
; GENERAL INFORMATION:
; APPLICANT: Beyer, Brian
; APPLICANT: Hammond, Gerald S
; APPLICANT: Reichert, Paul
; APPLICANT: Strickland, Corey
```

```

; APPLICANT: Wang, Wenyan
; APPLICANT: Weber, Patricia C
; APPLICANT: Wong, Gwendolyn
; APPLICANT: Zhang, Lili
; TITLE OF INVENTION: BETA-SECRETASE CRYSTALS AND METHODS FOR PREPARING AND USING THE S
; FILE REFERENCE: JB01531-K-US
; CURRENT APPLICATION NUMBER: US/10/400,273
; CURRENT FILING DATE: 2003-03-26
; PRIOR APPLICATION NUMBER: 60/367,937
; PRIOR FILING DATE: 2002-03-27
; NUMBER OF SEQ ID NOS: 5
; SOFTWARE: PatentIn version 3.1
; SEQ ID NO 5
; LENGTH: 408
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-400-273-5

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Query Match          100.0%; Score 144; DB 15; Length 408;
Best Local Similarity 100.0%; Pred. No. 4.1e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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```

Qy 1 GYVEMTVGSPPTLNIIVDTGSSNFAV 28
    |||||
Db 32 GYVEMTVGSPPTLNIIVDTGSSNFAV 59

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RESULT 11
US-10-400-273-1
; Sequence 1, Application US/10400273
; Publication No. US2004001494A1
; GENERAL INFORMATION:
; APPLICANT: Beyer, Brian
; APPLICANT: Hammond, Gerald S
; APPLICANT: Reichert, Paul
; APPLICANT: Strickland, Corey
; APPLICANT: Wang, Wenyan
; APPLICANT: Weber, Patricia C
; APPLICANT: Wong, Gwendolyn
; APPLICANT: Zhang, Lili
; TITLE OF INVENTION: BETA-SECRETASE CRYSTALS AND METHODS FOR PREPARING AND USING THE S
; FILE REFERENCE: JB01531-K-US
; CURRENT APPLICATION NUMBER: US/10/400,273
; CURRENT FILING DATE: 2003-03-26
; PRIOR APPLICATION NUMBER: 60/367,937
; PRIOR FILING DATE: 2002-03-27
; NUMBER OF SEQ ID NOS: 5
; SOFTWARE: PatentIn version 3.1
; SEQ ID NO 1
; LENGTH: 411
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-400-273-1

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Query Match          100.0%; Score 144; DB 15; Length 411;
Best Local Similarity 100.0%; Pred. No. 4.1e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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```

Qy 1 GYVEMTVGSPPTLNIIVDTGSSNFAV 28
    |||||
Db 29 GYVEMTVGSPPTLNIIVDTGSSNFAV 56

```

```

RESULT 12
US-10-627-473-19
; Sequence 19, Application US/10627473
; Publication No. US20040096950A1
; GENERAL INFORMATION:
; APPLICANT: VUILLARD, LAURENT MICHEL MARIE
; APPLICANT: PATEL, SAHIL JOE
; APPLICANT: YON, JEFFREY ROLAND
; APPLICANT: CLAESBY, ANNE
; APPLICANT: HAMILTON, BRUCE JOHN

```

```

; APPLICANT: SHAH, ALEEM
; TITLE OF INVENTION: CRYSTAL STRUCTURE OF BETA SITE APP CLEAVING ENZYME
; FILE REFERENCE: 674553-2002.1
; CURRENT APPLICATION NUMBER: US/10/627,473
; CURRENT FILING DATE: 2003-07-25
; PRIOR APPLICATION NUMBER: 60/398,681
; PRIOR FILING DATE: 2002-07-26
; NUMBER OF SEQ ID NOS: 46
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 19
; LENGTH: 411
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-627-473-19

```

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Query Match          100.0%; Score 144; DB 15; Length 411;
Best Local Similarity 100.0%; Pred. No. 4.1e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

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```

Qy 1 GYVEMTVGSPPTLNIIVDTGSSNFAV 28
    |||||
Db 32 GYVEMTVGSPPTLNIIVDTGSSNFAV 59

```

```

RESULT 13
US-10-627-473-20
; Sequence 20, Application US/10627473
; Publication No. US20040096950A1
; GENERAL INFORMATION:
; APPLICANT: VUILLARD, LAURENT MICHEL MARIE
; APPLICANT: PATEL, SAHIL JOE
; APPLICANT: YON, JEFFREY ROLAND
; APPLICANT: CLAESBY, ANNE
; APPLICANT: HAMILTON, BRUCE JOHN
; APPLICANT: SHAH, ALEEM
; TITLE OF INVENTION: CRYSTAL STRUCTURE OF BETA SITE APP CLEAVING ENZYME
; FILE REFERENCE: 674553-2002.1
; CURRENT APPLICATION NUMBER: US/10/627,473
; CURRENT FILING DATE: 2003-07-25
; PRIOR APPLICATION NUMBER: 60/398,681
; PRIOR FILING DATE: 2002-07-26
; NUMBER OF SEQ ID NOS: 46
; SOFTWARE: PatentIn Ver. 2.1
; SEQ ID NO 20
; LENGTH: 411
; TYPE: PRT
; ORGANISM: Homo sapiens
US-10-627-473-20

```

```

Query Match          100.0%; Score 144; DB 15; Length 411;
Best Local Similarity 100.0%; Pred. No. 4.1e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

```

```

Qy 1 GYVEMTVGSPPTLNIIVDTGSSNFAV 28
    |||||
Db 32 GYVEMTVGSPPTLNIIVDTGSSNFAV 59

```

```

RESULT 14
US-10-281-092-9
; Sequence 9, Application US/10281092
; Publication No. US20040121947A1
; GENERAL INFORMATION:
; APPLICANT: Ghosh, Arun K.
; APPLICANT: Tang, Jordan J.N.
; APPLICANT: Bilcer, Geoffrey
; APPLICANT: Chang, Manpin
; APPLICANT: Hong, Lin
; APPLICANT: Koelsch, Gerald E.
; APPLICANT: Loy, Jeffrey A.
; APPLICANT: Turner, Robert T., III

```

APPLICANT: Devasumadrum, Thippeswamy  
TITLE OF INVENTION: COMPOUNDS WHICH INHIBIT BETA-SECRETASE  
FILE REFERENCE: 2932.1001-004  
CURRENT APPLICATION NUMBER: US/10/281,092  
CURRENT FILING DATE: 2002-10-23  
PRIOR APPLICATION NUMBER: US 10/032,818  
PRIOR FILING DATE: 2001-12-28  
PRIOR APPLICATION NUMBER: PCT US01/50826  
PRIOR FILING DATE: 2001-12-28  
PRIOR APPLICATION NUMBER: US 60/258,705  
PRIOR FILING DATE: 2000-12-28  
PRIOR APPLICATION NUMBER: US 60/275,756  
PRIOR FILING DATE: 2001-03-14  
PRIOR APPLICATION NUMBER: US 60/335,952  
PRIOR FILING DATE: 2001-10-23  
PRIOR APPLICATION NUMBER: US 60/333,545  
PRIOR FILING DATE: 2001-11-27  
PRIOR APPLICATION NUMBER: US 60/348,464  
PRIOR FILING DATE: 2002-01-14  
PRIOR APPLICATION NUMBER: US 60/348,615  
PRIOR FILING DATE: 2002-01-14  
PRIOR APPLICATION NUMBER: US 60/390,804  
PRIOR FILING DATE: 2002-06-20  
PRIOR APPLICATION NUMBER: US 60/397,557  
PRIOR FILING DATE: 2002-07-19  
Remaining Prior Application data removed - See File Wrapper or PALM.  
NUMBER OF SEQ ID NOS: 59  
SOFTWARE: FastSeq for Windows Version 4.0  
SEQ ID NO 9  
LENGTH: 414  
TYPE: PRT  
ORGANISM: Unknown  
FEATURE:  
OTHER INFORMATION: memapsin 2  
US-10-281-092-9

Query Match 100.0%; Score 144; DB 16; Length 414;  
Best Local Similarity 100.0%; Pred. No. 4.1e-13;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTNLINLVDTGSSNPAV 28  
DB 34 GYVEMTVGSPPTNLINLVDTGSSNPAV 61

RESULT 15  
US-10-627-473-21  
Sequence 21, Application US/10627473  
Publication No. US20040096950A1  
GENERAL INFORMATION:  
APPLICANT: VUILLARD, LAURENT MICHEL MARIE  
APPLICANT: PATEL, SAHIL JOE  
APPLICANT: YON, JEFFREY ROLAND  
APPLICANT: CLEASBY, ANNE  
APPLICANT: HAMILTON, BRUCE JOHN  
APPLICANT: SHAH, ALEEM  
TITLE OF INVENTION: CRYSTAL STRUCTURE OF BETA SITE APP CLEAVING ENZYME  
FILE REFERENCE: 674553-2002.1  
CURRENT APPLICATION NUMBER: US/10/627,473  
CURRENT FILING DATE: 2003-07-25  
PRIOR APPLICATION NUMBER: 60/398,681  
PRIOR FILING DATE: 2002-07-26  
NUMBER OF SEQ ID NOS: 46  
SOFTWARE: PatentIn Ver. 2.1  
SEQ ID NO 21  
LENGTH: 417  
TYPE: PRT  
ORGANISM: Homo sapiens  
US-10-627-473-21

Query Match 100.0%; Score 144; DB 15; Length 417;

Best Local Similarity 100.0%; Pred. No. 4.2e-13;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTNLINLVDTGSSNPAV 28  
DB 32 GYVEMTVGSPPTNLINLVDTGSSNPAV 59

Search completed: July 26, 2005, 16:51:25  
Job time : 154 secs

**This Page Blank (uspto)**



GenCore version 5.1.6  
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OM protein - protein search, using sw model

Run on: July 26, 2005, 16:38:10 ; Search time 39 Seconds  
(without alignments)  
69.079 Million cell updates/sec

Title: US-10-726-967A-52

Perfect score: 144

Sequence: 1 GYVEMTVGSPPTINILVDTGSSNFAV 28

Scoring table: BLOSUM62  
Gapop.10.0 , Gapext 0.5

Searched: 283416 seqs, 96216763 residues

Total number of hits satisfying chosen parameters: 283416

Minimum DB seq length: 0

Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database : PIR 79:\*

1: PIR1:\*

2: PIR2:\*

3: PIR3:\*

4: PIR4:\*

Pred. No. is the number of results predicted by chance to have a  
score greater than or equal to the score of the result being printed,  
and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	144	100.0	501	2	A59090 aspartic proteinase
2	91	63.2	709	2	T29692 hypothetical prote
3	90	62.5	396	2	T47207 aspartic proteinase
4	87	60.4	405	2	A25379 saccharopepsin (EC
5	83	57.6	398	2	T33383 hypothetical prote
6	83	57.6	406	1	REHUK renin (EC 3.4.23.1
7	82	56.9	383	2	UC7573 pepsinogen C - Alt
8	82	56.9	384	2	A39314 gastricsin (EC 3.4
9	82	56.9	389	2	JEO371 pepsin C (EC 3.4.2
10	82	56.9	400	2	I47099 renin (EC 3.4.23.1
11	82	56.9	401	1	REMSK renin (EC 3.4.23.1
12	82	56.9	402	1	REMSK renin (EC 3.4.23.1
13	82	56.9	506	2	S71591 aspartic proteinase
14	82	56.9	508	2	D85056 probable aspartic
15	82	56.9	509	2	S49349 cyprosin (EC 3.4.2
16	82	56.9	596	2	S57971 aspartic proteinase
17	81	56.2	387	2	A45117 hypothetical prote
18	81	56.2	433	2	E96649 aspartic prote
19	81	56.2	513	2	T09739 aspartic endopepti
20	81	56.2	537	2	S50344 aspergillopepsin h
21	80	55.6	506	2	F86253 hypothetical prote
22	80	55.6	506	2	T07915 probable aspartic
23	80	55.6	569	2	S64957 aspergillopepsin I
24	79	54.9	375	2	C96715 protein P4N2.8 (im
25	79	54.9	165	2	S61602 probable membrane
26	78	54.2	344	1	KHPGD cathepsin D (EC 3.
27	78	54.2	377	1	PEMOCJ gastricsin (EC 3.4
28	78	54.2	388	2	JC7246 pepsinogen C - com
29	78	54.2	388	2	A29937 gastricsin (EC 3.4

30	78	54.2	392	1	A24608 gastricsin (EC 3.4
31	78	54.2	398	2	I51185 cathepsin D (EC 3.
32	78	54.2	407	1	KHRTD cathepsin D (EC 3.
33	78	54.2	412	1	KHRTD cathepsin D (EC 3.
34	77	53.5	394	2	B43356 gastricsin (EC 3.4
35	76	52.8	391	2	A43356 cathepsin E (EC 3.
36	76	52.8	396	2	A34401 cathepsin E (EC 3.
37	76	52.8	508	2	S19697 aspartic proteinase
38	76	52.8	509	2	S66516 oryzaasin (EC 3.4.2
39	75	52.1	383	2	A41443 pepsin (EC 3.4.23.
40	75	52.1	410	1	KHMSD cathepsin D (EC 3.
41	75	52.1	428	2	S47096 cyprosin (EC 3.4.2
42	75	52.1	435	2	T45035 cyprosin (EC 3.4.2
43	75	52.1	474	2	T12049 hypothetical prote
44	74	51.4	151	2	T18478 hypothetical prote
45	74	51.4	189	2	T18480

#### ALIGNMENTS

RESULT 1  
A59090 aspartic proteinase (EC 3.4.23.-) BACE precursor - human  
N:Alternate names: beta-secretase; beta-site APP cleaving enzyme  
C:Species: Homo sapiens (man)  
C>Date: 29-Oct-1999 #sequence\_revision 29-Oct-1999 #text\_change 09-Jul-2004  
C:Accession: A59090  
C:Vassar, R.; Bennett, B.D.; Babu-Khan, S.; Kahn, S.; Mendiaz, E.A.; Denis, P.; Teplow, M.A.; Biere, A.L.; Curran, E.; Burgess, T.; Louis, J.C.; Collins, F.; Treanor, J.; Roge, Science 286, 735-741, 1999  
A>Title: beta-Secretase cleavage of Alzheimer's amyloid precursor protein by the transm  
A:Reference number: A59090; PMID:20002972; PMID:10531052  
A>Note: submitted to GenBank, September 1999  
A:Accession: A59090  
A:Status: not compared with conceptual translation  
A:Molecule type: mRNA  
A:Residues: 1-501 <VASS>  
A:Cross-references: UNIPROT:P56817; GB:AF190725; NID:G6118538; PIDN:AAF04142.1; PID:G61  
C:Genetics:  
A:Gene: BACE  
C:Superfamily: beta-secretase  
C:Keywords: Alzheimer's disease; aspartic proteinase; brain; glycoprotein; hydrolase; p  
F:1-21/Domain: signal sequence #status predicted <SIG>  
F:22-45/Domain: propeptide #status predicted <PRO>  
F:46-501/Product: acid proteinase BACE #status predicted <MAT>  
F:461-477/Domain: transmembrane #status predicted <TRN>  
F:93,289/Active site: Asp #status predicted  
F:153,172,223,354/Binding site: carbohydrate (asn) (covalent) #status predicted  
F:330-380/Disulfide bonds: #status predicted

Query Match 100.0%; Score 144; DB 2; Length 501;  
Best Local Similarity 100.0%; Pred. No. 7,3e-14;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTINILVDTGSSNFAV 28  
DB 74 GYVEMTVGSPPTINILVDTGSSNFAV 101

RESULT 2  
T29692 hypothetical protein T189.2 - Caenorhabditis elegans  
C:Species: Caenorhabditis elegans  
C>Date: 15-Oct-1999 #sequence\_revision 15-Oct-1999 #text\_change 15-Oct-1999  
C:Accession: T29692  
R:Du, Z.; Gattlung, S.  
A:Submitted to the EMBL Data Library, November 1995  
A:Description: The sequence of C. elegans cosmid T189.  
A:Reference number: Z20666  
A:Accession: T29692  
A:Status: preliminary; translated from GB/EMBL/DBJ  
A:Molecule type: DNA

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A:Residues: 1-709 <DUZ>
A:Cross-references: EMBL:U41746; PIDN:AAA6331.1; CESP:TI8H9.2
C:Genetics:
A:Gene: CESP:TI8H9.2
A:Introns: 4/3; 126/3; 161/3; 190/3; 239/3; 282/3; 401/1; 691/3
Query Match
Best Local Similarity 63.2%; Score 91; DB 2; Length 709;
Matches 16; Conservative 5; Mismatches 5; Indels 0; Gaps 0;
OY 1 GYVEMTVGSPPTQNLIVDTGSSNF 26
Db 39 GYVGVTVGSPPTQFVMTDGSNF 64
RESULT 3
T47207
aspartic proteinase (EC 3.4.23.-) [imported] - Neurospora crassa
C:Species: Neurospora crassa
C>Date: 20-Apr-2000 #sequence_revision 20-Apr-2000 #text_change 09-Jul-2004
C:Accession: T47207
R:Bowman, B.
Submitted to the EMBL Data Library, September 1995
A:Reference number: Z24391
A:Accession: T47207
A:Status: preliminary; translated from GB/EMBL/DBJ
A:Molecule type: DNA
A:Residues: 1-396 <BOW>
A:Cross-references: UNIPROT:Q01294; EMBL:U36471; PIDN:AAA79878.1
C:Genetics:
A:Gene: pep-4
A:Introns: 33/3; 85/1
C:Superfamily: pepsin
C:Keywords: aspartic proteinase; hydrolase
Query Match
Best Local Similarity 62.5%; Score 90; DB 2; Length 396;
Matches 15; Conservative 7; Mismatches 5; Indels 0; Gaps 0;
OY 2 YVEMTVGSPPTQNLIVDTGSSNF 28
Db 85 YFSEITIGTPQTFKVLDTGSSNLMV 111
RESULT 4
Saccharopepsin (EC 3.4.23.25) precursor - yeast (Saccharomyces cerevisiae)
N:Alternate names: protein P2585; protein YPL154c; proteinase A; Saccharomyces aspartic
C:Species: Saccharomyces cerevisiae
C>Date: 25-Oct-1987 #sequence_revision 25-Oct-1987 #text_change 16-Aug-2004
C:Accession: A25379; A25378; A24711; B42640; S65165; S69445; S71635; S28231; S28233
R:Woolford, C.A.; Daniels, L.B.; Pak, F.D.; Jones, E.W.; Van Ardeell, J.N.; Innis, M.A.
Mol. Cell. Biol. 6, 2500-2510, 1986
A:Title: The PEP4 gene encodes an aspartyl protease implicated in the posttranslational
A:Reference number: A25379; MUID:87064548; PMID:3537721
A:Accession: A25379
A:Molecule type: DNA
A:Residues: 1-405 <WOO>
A:Cross-references: UNIPROT:P07267; EMBL:M13358; NID:g172121; PIDN:AAB63975.1; PID:g1721
R:Ammerer, G.; Hunter, C.P.; Rothman, J.H.; Saari, G.C.; Valls, L.A.; Stevens, T.H.
Mol. Cell. Biol. 6, 2490-2499, 1986
A:Title: PEP4 gene of Saccharomyces cerevisiae encodes proteinase A, a vacuolar enzyme
A:Reference number: A25378; MUID:87064547; PMID:3023936
A:Accession: A25378
A:Molecule type: DNA
A:Residues: 1-405 <AMM>
A:Cross-references: GB:M1338; GB:M1332; NID:g172121; PIDN:AAB63975.1; PID:g172122
R:Dreyer, T.; Halkier, B.; Svendsen, I.; Ottesen, M.
Carlsberg Res. Commun. 51, 27-41, 1986
A:Title: Primary structure of the aspartic proteinase A from Saccharomyces cerevisiae.
A:Reference number: A24711
A:Accession: A24711
A:Molecule type: protein
```

```
A:Residues: 77-405 <DRE>
R:Roof, D.M.; Meluh, P.B.; Rose, M.D.
J. Cell Biol. 118, 95-108, 1992
A:Title: Kinesin-related proteins required for assembly of the mitotic spindle.
A:Reference number: A42640; MUID:92317166; PMID:1618910
A:Accession: B42640
A:Molecule type: DNA
A:Residues: 373-405 <ROO>
A:Cross-references: EMBL:Z11963; NID:g3852; PIDN:CAA78020.1; PID:g3853
A>Note: the authors did not translate the codons for residues 373, 374, and 375
R:Purnelle, B.; Coater, F.; Goffeau, A.
Submitted to the Protein Sequence Database, May 1996
A:Reference number: S65154
A:Accession: S65154
A:Molecule type: DNA
A:Residues: 1-405 <PUR>
A:Cross-references: EMBL:Z73510; NID:g1370327; PIDN:CAA97859.1; PID:g1370328; MIPS:YPL15
A:Experimental source: strain 6286C (AB972)
R:Purnelle, B.; Comblez, S.; Coater, F.; Naveau, F.; Goffeau, A.
Submitted to the EMBL Data Library, March 1996
A:Description: The sequence of 55 kb on the left arm of yeast chromosome XVI identifies
ogues to the human phosphotyrosyl phosphatase activator PRPA and a homologue to the plant
A:Reference number: S69428
A:Accession: S69445
A:Molecule type: DNA
A:Residues: 1-405 <PUM>
A:Cross-references: EMBL:X96770; NID:g1403537; PIDN:CAA5567.1; PID:g1403555
R:Wolff, A.M.; Din, N.; Lillke Petersen, J.G.
Yeast 12, 823-832, 1996
A:Title: Vacuolar and extracellular maturation of Saccharomyces cerevisiae proteinase A
A:Reference number: S71635; MUID:56437971; PMID:8840459
A:Accession: S71635
A:Molecule type: protein
A:Residues: 23-31; 68-86 <WOL>
C:Genetics:
A:Gene: SCD:PEP4; PRA1; PHO9
A:Cross-references: SGD:S0006075; MIPS:YPL154C
A:Map position: 16L
C:Function:
A:Description: responsible for degradation of internalized alpha-factor receptor and a-f
alkaline phosphatase, acid trehalase, and vacuolar exopolysphatase
C:Superfamily: Pepsin
C:Keywords: aspartic proteinase; hydrolase; yeast vacuole
F:1-22/Domain: signal sequence #status predicted <SIG>
F:23-67/Domain: propeptide #status experimental <PRO>
F:68-405/Product: saccharopepsin #status experimental <MAT>
Query Match
Best Local Similarity 60.4%; Score 87; DB 2; Length 405;
Matches 14; Conservative 7; Mismatches 6; Indels 0; Gaps 0;
OY 2 YVEMTVGSPPTQNLIVDTGSSNF 28
Db 91 YVDITLIGTPQTFKVLDTGSSNLMV 117
RESULT 5
T33383
hypothetical protein H22K11.1 - Caenorhabditis elegans
C:Species: Caenorhabditis elegans
C>Date: 29-Oct-1999 #sequence_revision 29-Oct-1999 #text_change 09-Jul-2004
C:Accession: T33383
R:Beck, C.; Wamsley, P.; Keppler, N.
Submitted to the EMBL Data Library, July 1998
A:Description: The sequence of C. elegans coemid H22K11.
A:Reference number: Z21333
A:Accession: T33383
A:Status: preliminary; translated from GB/EMBL/DBJ
A:Molecule type: DNA
A:Residues: 1-398 <BEC>
A:Cross-references: UNIPROT:P55956; EMBL:A077544; PIDN:AAC64617.1; GSPDB:GN00028; CESP:1
A:Experimental source: strain Bristol N2; clone H22K11
```



JC7573  
pepsinogen C - African clawed frog  
N:Alternate names: progastricsin  
C:Species: Xenopus laevis (African clawed frog)  
C>Date: 30-Jun-2001 #sequence\_revision 30-Jun-2001 #text\_change 09-Jul-2004  
C:Accession: JC7573; PC7118  
R:Ikuzawa, M.; Inokuchi, T.; Kobayashi, K.; Yasunasu, S.  
J:Biochem. 129, 147-153, 2001  
A:Title: Amphibian pepsinogens: Purification and characterization of Xenopus pepsinogens  
A:Reference number: JC7573; MUID:21064922; PMID:11134969  
A:Contents: Stomach  
A:Accession: JC7573  
A:Molecule type: mRNA  
A:Residues: 1-383 <IKU>  
A:Cross-references: UNIPROT:Q9DEC3; DDBJ:AB045379  
A:Accession: PC7118  
A:Molecule type: protein  
A:Residues: 17-68 <IK2>  
C:Comment: This protein is a zymogen for gastric aspartic proteinase, with pepsin-like  
C:Genetics:  
A:Gene: Pgc  
C:Superfamily: pepsin  
C:Keywords: stomach; zymogen

Query Match 56.9%; Score 82; DB 2; Length 383;  
Best Local Similarity 55.6%; Pred. No. 0.00015;  
Matches 15; Conservative 5; Mismatches 7; Indels 0; Gaps 0;

QY 2 YVEMTVGSPQTLNLTVDGSSNFAV 28  
Db 67 YVGEISIGTPQNFVLVFDTGSSNLWV 93

RESULT 8  
A39314  
gastricsin (EC 3.4.23.3) precursor - bullfrog  
C:Species: Rana catesbeiana (bullfrog)  
C>Date: 19-Jun-1992 #sequence\_revision 19-Jun-1992 #text\_change 16-Aug-2004  
C:Accession: A39314  
R:Yakabe, E.; Tanji, M.; Ichinose, M.; Goto, S.; Miki, K.; Kurokawa, K.; Ito, H.; Kageya  
J: Biol. Chem. 266, 22436-22443, 1991  
A:Title: Purification, characterization, and amino acid sequences of pepsinogens and pep  
A:Reference number: A39314; MUID:92042186; PMID:1939266  
A:Accession: A39314  
A:Status: preliminary  
A:Molecule type: mRNA  
A:Residues: 1-384 <YKA>  
A:Cross-references: UNIPROT:Q91322; GB:M73750; NID:G213687; PID:AAA49530.1; PID:G213688  
C:Superfamily: Pepsin  
C:Keywords: aspartic proteinase; hydrolase; protein digestion

Query Match 56.9%; Score 82; DB 2; Length 384;  
Best Local Similarity 55.6%; Pred. No. 0.00015;  
Matches 15; Conservative 5; Mismatches 7; Indels 0; Gaps 0;

QY 2 YVEMTVGSPQTLNLTVDGSSNFAV 28  
Db 67 YVGEISIGTPQNFVLVFDTGSSNLWV 93

RESULT 9  
J50371  
pepsin C (EC 3.4.23.-) precursor - chicken  
N:Alternate names: pepsinogen C  
C:Species: Gallus gallus (chicken)  
C>Date: 23-Jul-1999 #sequence\_revision 23-Jul-1999 #text\_change 09-Jul-2004  
C:Accession: J50371  
R:Sakamoto, N.; Saiga, H.; Yasugi, S.  
Biochem. Biophys. Res. Commun. 250, 420-424, 1998  
A:Title: Analysis of temporal expression pattern and cis-regulatory sequences of chicken  
A:Reference number: J50370; MUID:9840813; PMID:9753665  
A:Accession: J50371  
A:Status: preliminary

A:Molecule type: mRNA  
A:Residues: 1-389 <SKA>  
A:Cross-references: UNIPROT:Q9W643; UNIPROT:Q9PMK1  
C:Superfamily: pepsin  
C:Keywords: aspartic proteinase; hydrolase

Query Match 56.9%; Score 82; DB 2; Length 389;  
Best Local Similarity 55.6%; Pred. No. 0.00016;  
Matches 15; Conservative 5; Mismatches 7; Indels 0; Gaps 0;

QY 2 YVEMTVGSPQTLNLTVDGSSNFAV 28  
Db 73 YVGEISIGTPQNFVLVFDTGSSNLWV 99

RESULT 10  
I47099  
renin (EC 3.4.23.15) - sheep  
C:Species: Ovis orientalis aries, Ovis ammon aries (domestic sheep)  
C>Date: 15-Oct-1996 #sequence\_revision 15-Oct-1996 #text\_change 09-Jul-2004  
C:Accession: I47099  
R:Alfred, G.P.; Fu, P.; Crawford, R.J.; Fernley, R.T.  
J: Mol. Endocrinol. 8, 3-11, 1992  
A:Title: The sequence and tissue expression of ovine renin.  
A:Reference number: I47099; MUID:92181567; PMID:1543552  
A:Accession: I47099  
A:Status: preliminary; translated from GB/EMBL/DDBJ  
A:Molecule type: mRNA  
A:Residues: 1-400 <ALD>  
A:Cross-references: UNIPROT:P52115; GB:I43524; NID:9896317; PID:AAA69809.1; PID:989631  
C:Superfamily: pepsin  
C:Keywords: aspartic proteinase; hydrolase  
P:98,286/Active site: Asp #status predicted

Query Match 56.9%; Score 82; DB 2; Length 400;  
Best Local Similarity 51.9%; Pred. No. 0.00016;  
Matches 14; Conservative 6; Mismatches 7; Indels 0; Gaps 0;

QY 2 YVEMTVGSPQTLNLTVDGSSNFAV 28  
Db 80 YVGEIGTGPQTFKVIPTDGSANLWV 106

RESULT 11  
REM5S  
renin (EC 3.4.23.15) precursor, submandibular - mouse  
C:Species: Mus musculus (house mouse)  
C>Date: 15-Oct-1982 #sequence\_revision 17-Dec-1982 #text\_change 16-Aug-2004  
C:Accession: A93283; A93285; B93285; B22058; A00988  
R:Misono, K.S.; Chang, J.U.; Inagami, T.  
Proc. Natl. Acad. Sci. U.S.A. 79, 4858-4862, 1982  
A:Title: Amino acid sequence of mouse submaxillary gland renin.  
A:Reference number: A93283; MUID:83014991; PMID:6812055  
A:Accession: A93283  
A:Molecule type: protein  
A:Residues: 64-351;354-401 <MIS>  
A:Cross-references: UNIPROT:P00796  
R:Pantlher, J.U.; Foote, S.; Chambrud, B.; Stroberg, A.D.; Corvol, P.; Rougeon, F.  
Nature 298, 90-92, 1982  
A:Title: Complete amino acid sequence and maturation of the mouse submaxillary gland ren  
A:Reference number: A93285; MUID:82220074; PMID:6283373  
A:Accession: A93285  
A:Molecule type: mRNA  
A:Residues: 1-98, 'W', 100-194, 'LSRS', 199-394, 'V', 396-401 <PA1>  
A:Cross-references: GB:J00621; GB:V00845; NID:9200701; PID:AAA00050.1; PID:9200702  
A:Note: the authors translated codon ATG for residue 99 as Ile  
A:Accession: B93285  
A:Molecule type: protein  
A:Residues: 64-84;354-374 <PA2>  
R:Poel, M.; Liesch, J.M.  
J: Biol. Chem. 258, 9856-9860, 1983  
A:Title: Mouse submaxillary gland renin contains a noncovalently attached fatty acid.  
A:Reference number: A92439; MUID:83290909; PMID:6350284

A:Contents: annotation; fatty acid binding  
R:Panther, J.J.; Dreyfus, M.; Roux, D.T.L.; Rougeon, F.  
Proc. Natl. Acad. Sci. U.S.A. 81, 5489-5493, 1984  
A:Title: Mouse kidney and submaxillary gland renin genes differ in their 5' putative reg  
A:Reference number: A22058; MUID:84298161; PMID:6089205  
A:Accession: B22058  
A:Molecule type: DNA  
A:Residues: 1-29 <PAN>  
C:Comment: The enzyme contains a noncovalently attached fatty acid.  
C:Comment: Submandibular renin has catalytic and antigenic activities similar to renal r  
C:Comment: This renin is synthesized in the submandibular gland of males only.  
C:Genetics:  
A:Gene: REN2  
C:Superfamily: Peprin  
C:Keywords: aspartic proteinase; hydrolase; salivary gland; submandibular gland  
F:1-21/Domain: signal sequence #status predicted <SIG>  
F:22-63/Domain: activation peptide #status predicted <ACP>  
F:64-351/Product: renin, submandibular, heavy chain #status experimental <RSH>  
F:354-401/Product: renin, submandibular, light chain #status experimental <RSL>  
F:101,286/Active site: Asp #status experimental  
F:114-121,277-281,320-357/Diulfide bonds: #status predicted

Query Match 56.9%; Score 82; DB 1; Length 401;  
Best Local Similarity 51.9%; Pred. No. 0.00016;  
Matches 14; Conservative 6; Mismatches 7; Indels 0; Gaps 0;

QY 2 YVEMTVGSPPTNTLIVDTGSSNFAY 28  
DB 83 YVGEIGIGTPPQTFKVIPTGSSNLMV 109

RESULT 12  
REMSK  
renin (EC 3.4.23.15) precursor, renal - mouse  
N:Alternate names: angiotensin-forming enzyme; angiotensinogenase; renin 1  
C:Species: Mus musculus (house mouse)  
C:Date: 30-Jun-1987 #sequence revision 30-Jun-1987 #text change 16-Aug-2004  
C:Accession: A00989; S07636; A22766; A22058; I57576; A05137; JH0083  
R:Holm, I.; Ollio, R.; Panther, J.J.; Rougeon, F.  
EMBO J. 3, 557-562, 1984  
A:Title: Evolution of aspartyl proteases by gene duplication: the mouse renin gene is on  
A:Reference number: A00989; MUID:84182525; PMID:6370686  
A:Accession: A00989  
A:Molecule type: DNA  
A:Residues: 1-402 <HOL>  
A:Cross-references: UNIPROT:P06281; EMBL:X00850  
R:Kim, W.S.; Murakami, K.; Nakayama, K.  
Nucleic Acids Res. 17, 9480, 1989  
A:Title: Nucleotide sequence of a cDNA coding for mouse Ren1 preprorenin.  
A:Reference number: S07636; MUID:90067953; PMID:2685761  
A:Accession: S07636  
A:Molecule type: mRNA  
A:Residues: 1-402 <KIM>  
A:Cross-references: EMBL:X16642; NID:953930; PIDN:CAA34636.1; PID:953931  
R:Nulling, U.J.; But, D.W.; Winkler, J.D.; McTurk, P.; George, H.; Brammar, W.J.  
EMBO J. 1, 1461-1466, 1982  
A:Title: Molecular cloning of two distinct renin genes from the DBA/2 mouse.  
A:Reference number: A90968; MUID:84207899; PMID:6327270  
A:Accession: A22766  
A:Molecule type: mRNA  
A:Residues: 269-314, 'D', 316 <MTL>  
R:Panther, J.J.; Dreyfus, M.; Roux, D.T.L.; Rougeon, F.  
Proc. Natl. Acad. Sci. U.S.A. 81, 5489-5493, 1984  
A:Title: Mouse kidney and submaxillary gland renin genes differ in their 5' putative reg  
A:Reference number: A22058; MUID:84298161; PMID:6089205  
A:Accession: A22058  
A:Molecule type: DNA  
A:Residues: 1-30 <PAN>  
R:Field, L.J.; Philbrick, W.M.; Howles, P.N.; Dickinson, D.P.; McGowan, R.A.; Gross, K.M.  
Mol. Cell. Biol. 4, 2321-2331, 1984  
A:Title: Expression of tissue-specific Ren-1 and Ren-2 genes of mice: Comparative analys  
A:Reference number: I57576; MUID:85085936; PMID:6392850  
A:Accession: I57576

A>Status: preliminary; translated from GB/EMBL/DBU  
A:Molecule type: DNA  
A:Residues: 1-31 <RES>  
A:Cross-references: GB:X02800; NID:9200689; PIDN:AAA40044.1; PID:9200690  
C:Comment: The only known function of renal renin is to release angiotensin I from angio  
created sodium retention by the kidney.  
C:Comment: Renal renin is synthesized by the juxtaglomerular cells of the kidney in resp  
C:Genetics:  
A:Gene: Ren-1  
A:Introns: 31/2; 81/3; 123/1; 162/3; 228/2; 268/2; 316/3; 349/3  
C:Superfamily: Peprin  
C:Keywords: aspartic proteinase; blood pressure control; glycoprotein; hydrolase; kidney  
F:1-21/Domain: signal sequence #status predicted <SIG>  
F:22-64/Domain: propeptide #status predicted <PRO>  
F:65-402/Product: renin #status predicted <MAT>  
F:59,139,320/Binding site: carbohydrate (Asn) (covalent) #status predicted  
F:102,287/Active site: Asp #status predicted

Query Match 56.9%; Score 82; DB 1; Length 402;  
Best Local Similarity 51.9%; Pred. No. 0.00016;  
Matches 14; Conservative 6; Mismatches 7; Indels 0; Gaps 0;

QY 2 YVEMTVGSPPTNTLIVDTGSSNFAY 28  
DB 84 YVGEIGIGTPPQTFKVIPTGSSNLMV 110

RESULT 13  
S71591  
aspartic proteinase precursor, wound-induced - tomato  
C:Species: Lycopersicon esculentum (tomato)  
C:Date: 04-Feb-1998 #sequence revision 13-Feb-1998 #text change 09-Jul-2004  
C:Accession: S71591  
R:Schaller, A.; Ryan, C.A.  
Plant Mol. Biol. 31, 1073-1077, 1996  
A:Title: Molecular cloning of a tomato leaf cDNA encoding an aspartic protease, a system  
A:Reference number: S71591; MUID:97000919; PMID:8843949  
A:Accession: S71591  
A>Status: preliminary  
A:Molecule type: mRNA  
A:Residues: 1-506 <SCH>  
A:Cross-references: UNIPROT:Q40140; EMBL:L46681; NID:9951448; PIDN:ABA18280.1; PID:99514  
C:Superfamily: oryzasin; sapsin repeat homology

Query Match 56.9%; Score 82; DB 2; Length 506;  
Best Local Similarity 55.6%; Pred. No. 0.00021;  
Matches 15; Conservative 4; Mismatches 8; Indels 0; Gaps 0;

QY 2 YVEMTVGSPPTNTLIVDTGSSNFAY 28  
DB 83 YVGEIGIGTPPQTFKVIPTGSSNLMV 109

RESULT 14  
D85056  
probable aspartic proteinase [imported] - Arabidopsis thaliana  
C:Species: Arabidopsis thaliana (mouse-ear cress)  
C:Date: 16-Feb-2001 #sequence revision 16-Feb-2001 #text change 09-Jul-2004  
C:Accession: D85056  
R:anonymous, The European Union Arabidopsis Genome Sequencing Consortium, The Cold Sprir  
Nature 402, 769-777, 1999  
A:Title: Sequence and analysis of chromosome 4 of the plant Arabidopsis thaliana.  
A:Reference number: A85001; MUID:20083488; PMID:10617198  
A:Accession: D85056  
A>Status: preliminary  
A:Molecule type: DNA  
A:Residues: 1-508 <STO>  
A:Cross-references: UNIPROT:Q9XEC4; GB:NC\_001268; NID:97267203; PIDN:CAB77914.1; GSPDB:Q  
C:Genetics:  
A:Gene: Atg04460  
A:Map position: 4  
C:Superfamily: oryzasin; sapsin repeat homology

Query Match 56.9%; Score 82; DB 2; Length 508;  
 Best Local Similarity 54.2%; Pred. No. 0.00021;  
 Matches 13; Conservative 6; Mismatches 5; Indels 0; Gaps 0;

OY 2 YVEMTVGSPQPTLNILVDTGSSN 25  
 ||:|||||:|||||  
 Db 87 YGDIITIGTPQKFTVIFDTGSSN 110

## RESULT 15

S49349  
 CYP19A1 (EC 3.4.23.-) - cardoon  
 C:Species: Cynara cardunculus (cardoon)  
 C>Date: 16-Feb-1995 #sequence\_revision 12-May-1995 #text\_change 09-Jul-2004  
 C:Accession: S49349  
 R:Pietrzak, M.; Brodeur, P.; Pais, M.S.  
 submitted to the EMBL Data Library, September 1994  
 A:Reference number: S49349  
 A:Accession: S49349  
 A:Status: preliminary  
 A:Molecule type: mRNA  
 A:Residues: 1-509 <PIB>  
 A:Cross-references: UNIPROT:039476; EMBL:X81984; NID:9556818; PIDN:CA57510.1; PID:95568  
 C:Comment: The pair of saposin repeat homology domains tagged SAP1 and SAP2 represent a  
 C:Superfamily: oryzasin; saposin repeat homology  
 C:Keywords: aspartic proteinase; hydrolase  
 F:316-361/Domain: saposin repeat homology #status atypical <SAP1>  
 F:375-420/Domain: saposin repeat homology #status atypical <SAP2>  
 F:103,290/Active site: Asp #status predicted

Query Match 56.9%; Score 82; DB 2; Length 509;  
 Best Local Similarity 55.6%; Pred. No. 0.00021;  
 Matches 15; Conservative 4; Mismatches 8; Indels 0; Gaps 0;

OY 2 YVEMTVGSPQPTLNILVDTGSSNFAV 28  
 ||:|||||:|||||  
 Db 85 YGDIITIGTPQKFTVIFDTGSSNLMV 111

Search completed: July 26, 2005, 16:47:58  
 Job time : 40 secs

GenCore version 5.1.6  
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OM protein - protein search, using sw model

Run on: July 26, 2005, 16:37:25 ; Search time 167 Seconds

(without alignments)  
85.858 Million cell updates/sec

Title: US-10-726-967a-52

Perfect score: 144

Sequence: 1 GYVEMTVGSPQNTLNLVDTGSSNPAV 28

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 1612378 seqs, 512079187 residues

Total number of hits satisfying chosen parameters: 1612378

Minimum DB seq length: 0  
Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database : UniProt 03:\*  
1: uniprot\_sprot:\*  
2: uniprot\_trembl:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	144	100.0	127	2	076KP0
2	144	100.0	457	2	08CAF4
3	144	100.0	501	1	BAE1_HUMAN
4	144	100.0	501	1	BAE1_MOUSE
5	144	100.0	501	1	BAE1_RAT
6	144	100.0	501	2	081YC8
7	144	100.0	501	2	08BOY4
8	144	100.0	532	2	09JUS1
9	143	99.3	501	2	08C7R1
10	124	86.1	505	2	06NZT7
11	122	84.7	499	2	07T0Y2
12	120	83.3	499	2	06PB20
13	119	82.6	396	2	09NZL1
14	119	82.6	439	2	09NZV8
15	119	82.6	468	2	09NZL2
16	119	82.6	518	1	BAE2_HUMAN
17	116	80.6	514	2	06IEF5
18	116	80.6	514	2	08CSE9
19	116	80.6	514	2	08C793
20	116	80.6	514	2	09JUL8
21	100	69.4	443	2	08WQY9
22	93	64.6	423	2	08NZD4
23	91	63.2	396	2	06C080
24	90	62.5	396	1	CARP_NEUCR
25	90	62.5	396	2	07RVB0
26	90	62.5	409	2	06CRK3
27	89	61.8	173	2	096TV7
28	89	61.8	427	2	P91802
29	89	61.8	428	2	07JNB4
30	89	61.8	429	2	026515
31	87	60.4	405	1	CARP_YEAST

32	86	59.7	398	2	06H2K5	06H2K5 botrytis ci
33	86	59.7	400	2	086ZP8	086ZP8 paracoccidi
34	86	59.7	504	2	093XR0	093XR0 ipomoea bat
35	85	59.0	354	2	09GYX7	09GYX7 boophilus m
36	84	58.3	397	2	08NUS2	08NUS2 leptosphaer
37	84	58.3	401	2	06TMM6	06TMM6 xenopus lae
38	84	58.3	406	2	06DLW5	06DLW5 macaca muli
39	83	57.6	358	2	09FRW5	09FRW5 nepenthes a
40	83	57.6	398	1	ASP3_CAEEL	P55956 caenorhabdi
41	83	57.6	400	1	RENI_CALJA	09552 callithrix
42	83	57.6	403	2	06R5C2	06R5C2 homo sapien
43	83	57.6	406	1	RENI_HUMAN	P00797 homo sapien
44	83	57.6	406	1	RENI_PANTR	P60016 pan troglod
45	83	57.6	406	2	06DLJ0	06DLJ0 macaca faec

## ALIGNMENTS

RESULT 1	PRELIMINARY	PRT	127 AA.
ID 076KP0			
AC 076KP0			
DT 05-JUL-2004 (TREMBlrel. 27, Created)			
DT 05-JUL-2004 (TREMBlrel. 27, Last sequence update)			
DT 05-JUL-2004 (TREMBlrel. 27, Last annotation update)			
DE Beta-site APP cleaving enzyme isoform I-127.			
GN Name=BACE;			
OS Homo sapiens (Human).			
OC Eukaryota; Metazoa; Chordata; Vertebrata; Euteleostomi;			
OC Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.			
OX NCBI_Taxid=9606;			
RN [1]			
RP SEQUENCE FROM N.A.			
RA Tanahashi H.;			
RL Submitted (Aug-2002) to the EMBL/GenBank/DBJ databases.			
CC -1- SIMILARITY: Belongs to peptidase family A1.			
DR EMBL; AB089958; BAC81826.1; -			
DR HSSP; P00797; 1BBS.			
DR GO; GO:0009049; F:aspartic-type signal peptidase activity; IEA.			
DR GO; GO:0006508; P:proteolysis and peptidolysis; IEA.			
DR InterPro; IPR001461; Peptidase_A1.			
DR InterPro; IPR009119; Pept_A1_BACE.			
DR InterPro; IPR009120; Pept_A1_BACE.			
DR InterPro; IPR009007; Pept_Aspartic.			
DR InterPro; IPR001969; Pept_Asp_AS.			
DR Pfam; PF00026; Asp_1.			
DR PRINTS; PRO1815; BACEFAMILY.			
DR PROSITE; PS00141; ASP_PROTEASE; 1.			
KW Aspartyl protease; Hydrolase; Protease.			
SQ SEQUENCE 127 AA; 13939 MW; C657354CBE72DC4 CRC64;			
Query Match	100.0%;	Score 144;	DB 2; Length 127;
Best Local Similarity	100.0%;	Pred. No. 4.6e-14;	
Matches	28;	Conservative	0; Mismatches 0; Indels 0; Gaps 0;
QY	1 GYVEMTVGSPQNTLNLVDTGSSNPAV 28		
Db	74 GYVEMTVGSPQNTLNLVDTGSSNPAV 101		
RESULT 2			
ID 08CAF4			
AC 08CAF4	PRELIMINARY;	PRT;	467 AA.
DT 01-MAR-2003 (TREMBlrel. 23, Created)			
DT 01-MAR-2003 (TREMBlrel. 23, Last sequence update)			
DT 01-MAR-2004 (TREMBlrel. 26, Last annotation update)			
DE Mus musculus 0 day neonate cerebellum cDNA, RIKEN full-length enriched			
DE library, clone: C230031E16 product: beta-site APP cleaving enzyme, full			
DE insert sequence.			
GN Name=Bace1; Synonyms=Bace;			

OC Mus musculus (Mouse).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
 ON NCBI\_TaxID=10090;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RC MEDLINE=9279253; PubMed=10349636; DOI=10.1016/S0076-6879(99)03004-9;  
 RA Carninci P., Hayashizaki Y.;  
 RT "High-efficiency full-length cDNA cloning.";  
 RL Meth. Enzymol. 303:19-44(1999).  
 RN [2]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RC MEDLINE=21085660; PubMed=11217851; DOI=10.1038/35055500;  
 RA RIKEN FANTOM Consortium;  
 RT "Functional annotation of a full-length mouse cDNA collection.";  
 RL Nature 409:665-690(2001).  
 RN [3]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RA The FANTOM Consortium;  
 RT "Analysis of the mouse transcriptome based on functional annotation of  
 Nature 420:563-573(2002).  
 RN [4]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RC MEDLINE=20493374; PubMed=11042159; DOI=10.1101/gr.145100;  
 RA Carninci P., Shibata Y., Hayatsu N., Sugahara Y., Shibata K., Itoh M.,  
 Kono H., Okazaki Y., Murakami M., Hayashizaki Y.;  
 RT "Normalization and subtraction of cap-trapper-selected cDNAs to  
 prepare full-length cDNA libraries for rapid discovery of new genes.";  
 RL Genome Res. 10:1617-1630(2000).  
 RN [5]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RC MEDLINE=20503913; PubMed=11076861; DOI=10.1101/gr.152600;  
 RA Shibata K., Itoh M., Aizawa K., Nagaoka S., Sasaki N., Carninci P.,  
 Kono H., Akiyama J., Nishi K., Katsunai T., Tashiro H., Itoh M.,  
 Sumi N., Ishii Y., Nakamura S., Hazama M., Nishine T., Harada A.,  
 Yamamoto R., Matsumoto H., Sakaguchi S., Ikegami T., Kashiwagi K.,  
 Fujiwaka S., Inoue K., Ozawa Y., Izawa M., Ohara E., Wachihi M.,  
 Yoneda Y., Ishikawa T., Ozawa K., Tanaka T., Matsura S., Kawai J.,  
 Okazaki Y., Murakami M., Inoue Y., Kita A., Hayashizaki Y.;  
 RT "RIKEN integrated sequence analysis (RISA) system-384-format  
 sequencing pipeline with 384 multicapillary sequencer.";  
 RL Genome Res. 10:1757-1771(2000).  
 RN [6]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Cerebellum;  
 RA Adachi J., Aizawa K., Akimura T., Arakawa T., Bono H., Carninci P.,  
 Fukuda S., Furuno M., Hanagaki T., Hara A., Hashizume W.,  
 Hayashida K., Hayatsu N., Hiramoto K., Hirao T., Hirozane T.,  
 Hori F., Imotani K., Ishii Y., Itoh M., Kagawa I., Kanda K.,  
 Katoh H., Kawai J., Kojima Y., Kondo S., Kono H., Kouda M., Koya S.,  
 Kuribara C., Matsuyama T., Miyazaki A., Murita M., Nakamura M.,  
 Nishi K., Nomura K., Numazaki R., Ohno M., Ohsato N., Okazaki Y.,  
 Saio R., Satoh H., Sakai C., Sakai K., Sakazume N., Sano H.,  
 Sasaki D., Shibata K., Shingawa A., Shiraki T., Sogabe Y., Tagami M.,  
 Tagawa A., Takahashi F., Takaku-Akaiura S., Takeda Y., Tanaka T.,  
 Tomaru A., Toya T., Yasunishi A., Muramatsu M., Hayashizaki Y.;  
 RT Submitted (Apr-2002) to the EMBL/GenBank/DBJ databases.  
 CC -1 - SIMILARITY: Belongs to peptidase family A1.  
 DR EMBL: AK082317; BAC38462.1; -.  
 DR HSSP: P56817; IFRN.  
 DR MGD: MGI:1346542; Bace1.  
 DR GO: GO:0005768; C:cytosol; ISS.  
 DR GO: GO:0005615; C:extracellular space; TAS.  
 DR GO: GO:0005794; C:Golgi apparatus; ISS.  
 DR GO: GO:0016021; C:integral to membrane; ISS.  
 DR GO: GO:0004190; F:aspartic-type endopeptidase activity; ISS.

DR GO: GO:000435; P:beta-amyloid metabolism; ISS.  
 DR GO: GO:0006509; P:membrane protein ectodomain proteolysis; ISS.  
 DR InterPro: IPR001461; Peptidase A1.  
 DR InterPro: IPR009119; Pept\_A1\_BACE.  
 DR InterPro: IPR009120; Pept\_A1\_BACE1.  
 DR InterPro: IPR009007; Pept\_Asp\_AS.  
 DR PRINTS: PRO1816; BACE1.  
 DR PRINTS: PRO1815; BACEFAMILY.  
 DR PRINTS: PRO0792; PEPsin.  
 DR PROSITE: PS00141; ASP\_PROTEASE\_1.  
 KM Aspartyl protease, Hydroxylase, Protease.  
 SQ SEQUENCE 467 AA; 52063 MW; 31AB674FF1843652 CRC64;  
 Query Match 100.0%; Score 144; DB 2; Length 467;  
 Best Local Similarity 100.0%; Pred. No. 2, 2e-13;  
 Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 QY 1 GYVEMTVGSPPTNTLIVDTGSSNFAV 28  
 DB 74 GYVEMTVGSPPTNTLIVDTGSSNFAV 101  
 RESULT 3  
 BAE1\_HUMAN STANDARD; PRT; 501 AA.  
 AC P56817; Q9BYB9; Q9BYC0; Q9BYC1; Q9LUT5;  
 DT 30-MAY-2000 (Rel. 39, Created)  
 DT 30-MAY-2000 (Rel. 39, Last sequence update)  
 DT 25-OCT-2004 (Rel. 45, Last annotation update)  
 DE Beta-secretase 1 precursor (BC 3.4.23.46) (Beta-site APP cleaving  
 enzyme 1) (Beta-site amyloid precursor protein cleaving enzyme 1)  
 DE (Aspartyl protease 2) (Asp 2) (ASP2) (Membrane-associated aspartic  
 DE protease 2) (Memapsin-2).  
 GN Name=BACE1; Synonyms=BACE;  
 OS Homo sapiens (Human).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.  
 ON NCBI\_TaxID=9606;  
 RN [1]  
 RP SEQUENCE FROM N.A. (ISOFORM A).  
 RC TISSUE=Brain;  
 RC MEDLINE=20002972; PubMed=10511052; DOI=10.1126/science.286.5440.735;  
 RA Vassar R., Bennett B.D., Babu-Khan S., Kahn S., Mendiaz E.A.,  
 Denis P., Teplow D.B., Ross S., Amarante P., Loisel R., Luo Y.,  
 Fisher S., Fuller J., Edenson S., Lile J., Jarosinski M.A.,  
 Biere A.L., Curran E., Burgess T., Louis J.C., Collins F.,  
 Treanor J., Rogers G., Citron M.;  
 RT "Beta-secretase cleavage of Alzheimer's amyloid precursor protein by  
 the transmembrane aspartic protease BACE.";  
 RL Science 286:735-741(1999).  
 RN [2]  
 RP SEQUENCE FROM N.A. (ISOFORM A), SEQUENCE OF 46-68, AND  
 RP CHARACTERIZATION.  
 RC TISSUE=Brain;  
 RC MEDLINE=20057171; PubMed=10591214; DOI=10.1038/990114;  
 RA Sinha S., Anderson J.P., Barbour R., Basl G.S., Caccavello R.,  
 Davis D., Dean M., Dovey H.F., Frigon N., Hong J., Jacobson-Croak K.,  
 Jewett N., Keim P., Knops J., Lieberburg I., Fowler M., Tan H.,  
 Tetsuno G., Tung J., Schenk D., Seubert P., Suenematsu S.M., Wang S.,  
 Walker D., Zhao J., McConlogue L., Varghese J.;  
 RT "Purification and cloning of amyloid precursor protein beta-secretase  
 from human brain.";  
 RL Nature 402:537-540(1999).  
 RN [3]  
 RP SEQUENCE FROM N.A. (ISOFORM A)  
 RC MEDLINE=20057170; PubMed=10591213; DOI=10.1038/990107;  
 RA Van R., Bienkowski M.J., Snuck M.E., Miao H., Tory M.C., Pauley A.M.,  
 Braehler J.R., Stratman N.C., Mathews W.R., Buhl A.E., Carter D.B.,  
 Tomasekelli A.G., Parodi L.A., Heinrichson R.L., Guney M.E.;  
 RT "Membrane-anchored aspartyl protease with Alzheimer's disease beta-  
 RT secretase activity.";  
 RL Nature 402:533-537(1999).



[4] SEQUENCE FROM N.A. (ISOFORM A).  
 MEDLINE=20120043; PubMed=10656250; DOI=10.1006/mcne.1999.0811;  
 RA Husain I., Powell D.J., Howlett D.R., Tew D.G., Meek T.D.,  
 RA Chapman C., Gloger I.S., Murphy K.E., Southan C.D., Ryan D.M.,  
 RA Smith T.S., Simmons D.L., Walsh F.S., Dingwall C., Christie G.;  
 RT "Identification of a novel aspartic proteinase (Asp 2) as beta-  
 RT secretase";  
 RL Mol. Cell. Neurosci. 14:419-427(1999).  
 [5] SEQUENCE FROM N.A. (ISOFORM B).  
 TISSUE=Brain, and Pancreas;  
 RA Michel B., De Pietri Tonelli D., Zaccchetti D., Keller P.;  
 RT "New beta-site APP cleaving enzyme isoform (BACE-1B) obtained from  
 RT human brain and pancreas";  
 RL Submitted (JAN-2001) to the EMBL/GenBank/DBJ databases.  
 [6] SEQUENCE FROM N.A. (ISOFORM C).  
 TISSUE=Pancreas;  
 RA Zaccchetti D., De Pietri Tonelli D., Schnurbe R.;  
 RT "New beta-site APP cleaving enzyme isoform (BACE-1C) obtained from  
 RT human pancreas";  
 RL Submitted (JAN-2001) to the EMBL/GenBank/DBJ databases.  
 [7] SEQUENCE FROM N.A. (ISOFORMS B; C AND D).  
 TISSUE=Brain;  
 RA MEDLINE=21408467; PubMed=11516562; DOI=10.1016/S0304-3940(01)01912-7;  
 RA Tanahashi H., Tabira T.;  
 RT "Three novel alternatively spliced isoforms of the human beta-site  
 RT amyloid precursor protein cleaving enzyme (BACE) and their effect on  
 RT amyloid beta-peptide production";  
 RL Neurosci. Lett. 307:9-12(2001).  
 [8] SEQUENCE OF 14-501 FROM N.A. (ISOFORM A), AND CHARACTERIZATION.  
 RX MEDLINE=20144060; PubMed=10677483; DOI=10.1073/pnas.97.4.1456;  
 RA Lin X., Koelsch G., Wu S., Downs D., Dashedi A., Tang J.;  
 RT "Human aspartic protease memapsin 2 cleaves the beta-secretase site of  
 RT beta-amyloid precursor protein";  
 RL Proc. Natl. Acad. Sci. U.S.A. 97:1456-1460(2000).  
 [9] DISULFIDE BONDS.  
 RX MEDLINE=21950860; PubMed=11953458;  
 RA Fischer F., Molinari M., Bodendorf U., Paganetti P.;  
 RT "The disulphide bonds in the catalytic domain of BACE are critical but  
 RT not essential for amyloid precursor protein processing activity";  
 RL J. Neurochem. 80:1079-1088(2002).  
 -1- FUNCTION: Responsible for the proteolytic processing of the  
 CC amyloid precursor protein (APP). Cleaves at the amino terminus of  
 CC the A-beta peptide sequence, between residues 671 and 672 of APP,  
 CC leads to the generation and extracellular release of beta-cleaved  
 CC soluble APP, and a corresponding cell-associated carboxy-terminal  
 CC fragment which is later released by gamma-secretase.  
 -1- CATALYTIC ACTIVITY: Broad endopeptidase specificity. Cleaves Glu-  
 CC Val-Asn-Ileu-| Asp-Ala-Glu-Phe in the Swedish variant of  
 CC Alzheimer's amyloid precursor protein.  
 CC -1- SUBCELLULAR LOCATION: Type I membrane protein.  
 CC -1- ALTERNATIVE PRODUCTS:  
 CC Event=Alternative splicing; Named isoforms=4;  
 CC Name=A; Synonyms=BACE-1A, BAC-501;  
 CC IsoId=P56817-1; Sequence=Displayed;  
 CC Name=B; Synonyms=BACE-1B, BACE-1-476;  
 CC IsoId=P56817-2; Sequence=VSP\_005223;  
 CC Name=C; Synonyms=BACE-1C, BACE-1-457;  
 CC IsoId=P56817-3; Sequence=VSP\_005222;  
 CC Name=D; Synonyms=BACE-1D, BACE-1-432;  
 CC IsoId=P56817-4; Sequence=VSP\_005222, VSP\_005223;  
 CC -1- TISSUE SPECIFICITY: Brain.  
 CC -1- SIMILARITY: Belongs to the peptidase A1 family.  
 CC -----  
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 CC EMBL; AF190725; AAF04142.1; -  
 DR EMBL; AF201468; AAF18982.1; -  
 DR EMBL; AF200343; AAF17079.1; -  
 DR EMBL; AF204943; AAF26367.1; -  
 DR EMBL; AF338816; AAK38374.1; -  
 DR EMBL; AF338817; AAK38375.1; -  
 DR EMBL; AB050436; BAB40931.1; -  
 DR EMBL; AB050437; BAB40932.1; -  
 DR EMBL; AB050438; BAB40933.1; -  
 DR EMBL; AF200193; AAF13715.1; -  
 DR PIR; A59090; A59090.  
 DR PDB; 1FXN; X-ray; A/B=56-446.  
 DR PDB; 1M4H; X-ray; A/B=56-446.  
 DR MEROPS; A01.004; -  
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 DR MIM; 604252; -  
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 DR InterPro; IPR001969; Pept\_Asp\_AS.  
 DR InterPro; IPR001461; Peptidase\_A1.  
 DR Pfam; PF00026; Asp; 1.  
 DR PRINTS; PRO1816; BACE1.  
 DR PRINTS; PRO0792; BACEFAMILY.  
 DR PRINTS; PRO0792; BPSIN.  
 DR PROSITE; PS00141; ASP\_PROTEASE; 1.  
 DR 3D-structure; Alternative splicing; Aspartyl protease;  
 KW Direct protein sequencing; Glycoprotein; Hydrolase; Signal;  
 KW Transmembrane; Zymogen.  
 FT PROPR 1 21  
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 FT CHAIN 22 45  
 FT DOMAIN 22 457  
 FT TRANSMEM 458 478  
 FT DOMAIN 479 501  
 FT ACT\_SITE 93 93  
 FT ACT\_SITE 289 289  
 FT DISULFID 216 420  
 FT DISULFID 278 443  
 FT DISULFID 330 380  
 FT CARBOHYD 153 153  
 FT CARBOHYD 172 172  
 FT CARBOHYD 223 223  
 FT CARBOHYD 354 354  
 FT VARSPPLIC 146 189  
 FT VARSPPLIC 190 214  
 FT HELIX 61 63  
 FT TURN 64 65  
 FT STRAND 67 70  
 FT TURN 71 73  
 FT STRAND 74 81  
 FT TURN 82 85  
 FT STRAND 86 93  
 FT TURN 94 95  
 FT STRAND 99 102  
 FT TURN 107 108  
 FT TURN 115 117  
 FT HELIX 119 120  
 FT TURN 122 131  
 FT STRAND 136 147  
 FT TURN 149 150  
 FT STRAND 155 167  
 FT TURN 172 173  
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 N-linked (GlcNAc. . .) (Potential).  
 N-linked (GlcNAc. . .) (Potential).  
 N-linked (GlcNAc. . .) (Potential).  
 N-linked (GlcNAc. . .) (Potential).  
 Missing (in isoform C and isoform D).  
 /FtId=VSP\_005222.  
 Missing (in isoform B and isoform D).  
 /FtId=VSP\_005223.



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DR InterPro: IPR001461; Peptidase_A1.
DR Pfam: PF00026; Asp. 1.
DR PRINTS; PRO1816; BACE1.
DR PRINTS; PRO1815; BACEFAMILY.
DR PRINTS; PRO0792; PEPsin.
DR PROSITE; PS00141; ASP_PROTEASE; 1.
KW Aspartyl protease; Glycoprotein; Hydrolase; Signal; Transmembrane;
KM Zymogen.
FT SIGNAL 1 21 Potential.
FT PROPEP 22 45 Potential.
FT CHAIN 46 501 Beta-secretase 1.
FT DOMAIN 22 457 Extracellular (Potential).
FT TRANSMEM 458 478 Potential.
FT DOMAIN 479 501 Cytoplasmic (Potential).
FT ACT_SITE 93 93 By similarity.
FT ACT_SITE 289 289 By similarity.
FT DISULFID 216 420 By similarity.
FT DISULFID 278 443 By similarity.
FT DISULFID 330 380 By similarity.
FT CARBOHYD 153 153 N-linked (GlcNAc...) (Potential).
FT CARBOHYD 172 172 N-linked (GlcNAc...) (Potential).
FT CARBOHYD 223 223 N-linked (GlcNAc...) (Potential).
FT CARBOHYD 354 354 N-linked (GlcNAc...) (Potential).
SQ SEQUENCE 501 AA; 55747 MW; C085A013145E474E CRC64;

Query Match 100.0%; Score 144; DB 1; Length 501;
Best Local Similarity 100.0%; Pred. No. 2,4e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTQTLNILDVTSNFAV 28
Db 74 GYVEMTVGSPPTQTLNILDVTSNFAV 101

RESULT 5
ID BAE1_RAT STANDARD; PRT; 501 AA.
AC P56819;
DT 30-MAY-2000 (Rel. 39, Last sequence update)
DT 25-OCT-2004 (Rel. 45, Last annotation update)
DE Beta-secretase 1 precursor (EC 3.4.23.46) (Beta-site APP cleaving
enzyme 1) (beta-site amyloid precursor protein cleaving enzyme 1)
DE (Aspartyl protease 2) (Asp 2) (ASp2) (Membrane-associated aspartic
protease 2) (Memapsin-2).
GN Name=BACE1; Synonyms=Bace;
OS Rattus norvegicus (Rat).
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Rodentia; Scturognathi; Muridae; Murinae; Rattus.
OX NCBI_TaxID=10116;
RN [1]
RP SEQUENCE FROM N.A.
RX MEDLINE=20002972; Pubmed=10531052; DOI=10.1126/science.286.5440.735;
RA Vassar R., Bennett B.D., Babu-Khan S., Kahn S., Mendiaz E.A.,
RA Denis P., Teplow D.B., Ross S., Amarante P., Loebner R., Luo Y.,
RA Fisher S., Fuller J., Edenson S., Lile J., Jarosinski M.A.,
RA Biere A.L., Curran E., Burgess T., Louis J.-C., Collins F.,
RA Treanor J., Rogers G., Citron M.;
RA "Beta-secretase cleavage of Alzheimer's amyloid precursor protein by
the transmembrane aspartic protease BACE.";
RT Science 286:735-741(1999).
RL
CC -1- FUNCTION: Responsible for the proteolytic processing of the
CC amyloid precursor protein (APP). Cleaves at the amino terminus of
CC the A-beta peptide sequence, between residues 671 and 672 of APP,
CC leads to the generation and extracellular release of beta-cleaved
CC soluble APP, and a corresponding cell-associated carboxy-terminal
CC fragment which is later released by gamma-secretase (By
CC similarity).
CC -1- CATALYTIC ACTIVITY: Broad endopeptidase specificity. Cleaves Glu-
CC Val-Ileu-Leu-Ileu-Ala-Glu-Phe in the Swedish variant of
CC Alzheimer's amyloid precursor protein.
CC -1- SUBCELLULAR LOCATION: Type I membrane protein.
CC -1- SIMILARITY: Belongs to the peptidase A1 family.

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CC or send an email to license@sib-sib.ch).
CC -----
DR EMBL; AF190727; AAF04144.1; -.
DR HSSP; P56817; 1MAH.
DR MEROPS; A01.004; -.
DR RGD; 2191; Bace.
DR InterPro; IPR009119; Pept_A1_BACE.
DR InterPro; IPR009120; Pept_A1_BACE1.
DR InterPro; IPR001969; Pept_Asp_AS.
DR InterPro; IPR009007; Pept_AspArtic.
DR InterPro; IPR001461; Peptidase_A1.
DR Pfam; PF00026; Asp. 1.
DR PRINTS; PRO1816; BACE1.
DR PRINTS; PRO1815; BACEFAMILY.
DR PROSITE; PS00141; ASP_PROTEASE; 1.
DR PROSITE; PS00792; PEPsin.
KW Aspartyl protease; Glycoprotein; Hydrolase; Signal; Transmembrane;
KM Zymogen.
FT SIGNAL 1 21 Potential.
FT PROPEP 22 45 Potential.
FT CHAIN 46 501 Beta-secretase 1.
FT DOMAIN 22 457 Extracellular (Potential).
FT TRANSMEM 458 478 Potential.
FT DOMAIN 479 501 Cytoplasmic (Potential).
FT ACT_SITE 93 93 By similarity.
FT ACT_SITE 289 289 By similarity.
FT DISULFID 216 420 By similarity.
FT DISULFID 278 443 By similarity.
FT DISULFID 330 380 By similarity.
FT CARBOHYD 153 153 N-linked (GlcNAc...) (Potential).
FT CARBOHYD 172 172 N-linked (GlcNAc...) (Potential).
FT CARBOHYD 223 223 N-linked (GlcNAc...) (Potential).
FT CARBOHYD 354 354 N-linked (GlcNAc...) (Potential).
SQ SEQUENCE 501 AA; 55806 MW; 24B445BC8B87DE3 CRC64;

Query Match 100.0%; Score 144; DB 1; Length 501;
Best Local Similarity 100.0%; Pred. No. 2,4e-13;
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTQTLNILDVTSNFAV 28
Db 74 GYVEMTVGSPPTQTLNILDVTSNFAV 101

RESULT 6
ID Q81YC8 PRELIMINARY; PRT; 501 AA.
AC Q81YC8;
DT 01-MAR-2003 (TrEMBLrel. 23, Created)
DT 01-MAR-2003 (TrEMBLrel. 23, Last sequence update)
DT 01-MAR-2004 (TrEMBLrel. 26, Last annotation update)
DE Beta-site APP-cleaving enzyme 1, isoform A preproprotein.
DE Name=BACE1;
GN Homo sapiens (Human).
OS Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.
OX NCBI_TaxID=9606;
RN [1]
RP SEQUENCE FROM N.A.
RX MEDLINE=22386257; Pubmed=12477932; DOI=10.1073/pnas.242603899;
RA Strausberg R.L., Feingold E.A., Grouse L.H., Derge J.G.,
RA Klausner R.D., Collins F.S., Wagner L., Shenmen C.M., Schuler G.D.,
RA Altshuler S.F., Zeeberg B., Buetow K.H., Schaefer C.F., Bhat N.K.,
RA Hopkins R.F., Jordan H., Moore T., Max S.I., Wang J., Hsieh F.,
RA Datchenko L., Marusina K., Farmer A.A., Rubin G.M., Hong L.,

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RA Stagleton M., Soares M.B., Bonaldo M.F., Casavant T.L., Scheetz T.E.,  
 RA Brownstein M.J., Uedin T.B., Toshiyuki S., Carninci P., Prange C.C.,  
 RA Rana S.S., Loquellano N.A., Peters G.J., Abrahamson R.D., Mullahy S.J.,  
 RA Bosak S.A., McEwan P.J., McKernan K.J., Malek J.A., Gunatane P.H.,  
 RA Richards S., Morley K.C., Hale S., Garcia A.M., Gay L.J., Huilyk S.W.,  
 RA Villalón D.K., Muzny D.M., Sodergren E.J., Lu X., Gibbs R.A.,  
 RA Fahey J., Helton E., Kettelman M., Madan A., Rodrigues S., Sanchez A.,  
 RA Whiting M., Madan A., Young A.C., Shevchenko Y., Bouffard G.C.,  
 RA Blakeley R.W., Touchman J.W., Green E.D., Dickson M.C.,  
 RA Rodriguez A.C., Grimwood J., Schmutz J., Myers R.M., Butterfield Y.S.,  
 RA Krywinski M.I., Skaleka U., Smallus D.E., Scherren A., Schein J.E.,  
 RA Jones S.J., Maira M.A.,  
 RT "Generation and initial analysis of more than 15,000 full-length human  
 RT and mouse cDNA sequences."  
 RL Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903(2002).  
 RN [2]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Brain;  
 RL Submitted (JUL-2002) to the EMBL/GenBank/DBJ databases.  
 CC -1- SIMILARITY: Belongs to peptidase family A1.  
 DR EMBL; BC036084; AA036084.1; -.  
 DR HSSP; P56817; 1PKN.  
 DR GO; GO:0005768; C:endosome; ISS.  
 DR GO; GO:0005794; C:Golgi apparatus; ISS.  
 DR GO; GO:0016021; C:integral to membrane; ISS.  
 DR GO; GO:0004190; F:aspartic-type endopeptidase activity; ISS.  
 DR GO; GO:0005043; P:beta-amyloid metabolism; ISS.  
 DR GO; GO:0006509; P:membrane protein ectodomain proteolysis; ISS.  
 DR InterPro; IPR001461; Peptidase A1.  
 DR InterPro; IPR009119; Pept\_A1\_BACE.  
 DR InterPro; IPR009120; Pept\_A1\_BACE.  
 DR InterPro; IPR009007; Pept\_Aspartic.  
 DR InterPro; IPR001969; Pept\_Asp\_AS.  
 DR PRINTS; PRO1816; BACEFAMILY.  
 DR PRINTS; PRO1815; BACEFAMILY.  
 DR PRINTS; PRO0792; PEPsin.  
 DR PROSITE; PS00141; ASP\_PROTEASE; 1.  
 KM Aspartyl protease; Hydrolase; Protease.  
 SQ SEQUENCE 501 AA; 55823 MW; 7685955C5517EFB7 CRC64;  
 QY 1 GYVENTVSSPPTLITLVDTGSSNFAV 28  
 Db 74 GYVENTVSSPPTLITLVDTGSSNFAV 101  
 RESULT 7  
 O8BOY4 PRELIMINARY; PRT; 501 AA.  
 ID O8BOY4;  
 AC O8BOY4;  
 DT 01-MAR-2003 (TREMBLrel. 23, Created)  
 DT 01-MAR-2003 (TREMBLrel. 23, Last sequence update)  
 DT 01-MAR-2004 (TREMBLrel. 26, Last annotation update)  
 DE Mus musculus adult male corpora quadrigemina cDNA, RIKEN full-length  
 DE enriched library, clone:B230346M13 product:beta-site APP cleaving  
 DE enzyme, full insert sequence.  
 GN Name=Bace1; Synonym=Bace;  
 OS Mus musculus (Mouse).  
 OC Eukaryota; Euteleostomi; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
 OX NCBI\_TaxId=10090;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RX MEDLINE=99279253; PubMed=10349636; DOI=10.1016/S0076-6879(99)03004-9;  
 RA Carninci P., Hayashizaki Y.;  
 RT "High-efficiency full-length cDNA cloning."  
 RL Meth. Enzymol. 303:19-44(1999).  
 RN [2]

RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RX MEDLINE=21085660; PubMed=11217851; DOI=10.1038/35055500;  
 RA RIKEN FANTOM Consortium;  
 RT "Functional annotation of a full-length mouse cDNA collection."  
 RL Nature 409:685-690(2001).  
 RN [3]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RA The FANTOM Consortium,  
 RT "Analysis of the mouse transcriptome based on functional annotation of  
 RT 60,770 full-length cDNAs."  
 RL Nature 420:563-573(2002).  
 RN [4]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RX MEDLINE=20499374; PubMed=11042159; DOI=10.1101/gr.145100.  
 RA Carninci P., Shibata Y., Hayatsu N., Sugahara Y., Shibata K., Itoh M.,  
 RA Kono H., Okazaki Y., Muramatsu M., Hayashizaki Y.;  
 RT "Normalization and subcloning of cap-trapper-selected cDNAs to  
 RT prepare full-length cDNA libraries for rapid discovery of new genes."  
 RL Genome Res. 10:1617-1630(2000).  
 RN [5]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RX MEDLINE=20530913; PubMed=11076861; DOI=10.1101/gr.152600;  
 RA Shibata K., Itoh M., Aizawa K., Nagaoka S., Sasaki N., Carninci P.,  
 RA Kono H., Akiyama Y., Nishi K., Kitanai T., Tashiro H., Itoh M.,  
 RA Suni N., Ishii Y., Nakamura S., Hazama M., Nishine T., Harada A.,  
 RA Yamamoto R., Matsumoto H., Sakaguchi S., Ikegami T., Kashiwagi K.,  
 RA Fujiwara S., Inoue K., Togawa Y., Izawa M., Ohara E., Watabiki M.,  
 RA Yoneda Y., Ishikawa T., Ozawa K., Tanaka T., Matsumura S., Kawai J.,  
 RA "RIKEN integrated sequence analysis (RISA) system-384-format  
 RT sequencing pipeline with 384 multicapillary sequencer."  
 RL Genome Res. 10:1757-1771(2000).  
 RN [6]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Corpora quadrigemina;  
 RA Adachi J., Aizawa K., Akimura T., Arikawa T., Bono H., Carninci P.,  
 RA Fukuda S., Furuno M., Hanagaki T., Hara A., Hashizume W.,  
 RA Hayashida K., Hayatsu N., Hiramoto K., Hiraoka T.,  
 RA Hori F., Imotani K., Ishii Y., Itoh M., Kagawa I., Kasukawa T.,  
 RA Katoh H., Kawai J., Kojima Y., Kondo S., Kono H., Kouda M., Koya S.,  
 RA Kurihara C., Matsuyama T., Miyazaki A., Murata M., Nakamura M.,  
 RA Nishi K., Nomura K., Numazaki R., Ono M., Ono S., Okazaki Y.,  
 RA Saito R., Saitoh H., Sakai C., Sakai K., Sakazume N., Sano H.,  
 RA Sasaki D., Shibata K., Shinagawa A., Shiraki T., Sogabe Y., Tagami M.,  
 RA Tomaru A., Toyota T., Yasunishi A., Muramatsu M., Hayashizaki Y.,  
 RA Submitted (JUL-2001) to the EMBL/GenBank/DBJ databases.  
 CC -1- SIMILARITY: Belongs to peptidase family A1.  
 DR EMBL; AK046175; BAC32820.1; -.  
 DR HSSP; P56817; 1PKN.  
 DR MGI; MGI:1346542; Bace1.  
 DR GO; GO:0005768; C:endosome; ISS.  
 DR GO; GO:0005615; C:extracellular space; TNS.  
 DR GO; GO:0005794; C:Golgi apparatus; ISS.  
 DR GO; GO:0016021; C:integral to membrane; ISS.  
 DR GO; GO:0004190; F:aspartic-type endopeptidase activity; ISS.  
 DR GO; GO:0005043; P:beta-amyloid metabolism; ISS.  
 DR GO; GO:0006509; P:membrane protein ectodomain proteolysis; ISS.  
 DR InterPro; IPR001461; Peptidase A1.  
 DR InterPro; IPR009119; Pept\_A1\_BACE.  
 DR InterPro; IPR009120; Pept\_A1\_BACE.  
 DR InterPro; IPR009007; Pept\_Aspartic.  
 DR InterPro; IPR001969; Pept\_Asp\_AS.  
 DR PRINTS; PRO1816; BACE1.  
 DR PRINTS; PRO1815; BACEFAMILY.  
 DR PRINTS; PRO0792; PEPsin.  
 DR PROSITE; PS00141; ASP\_PROTEASE; 1.  
 KM Aspartyl protease; Hydrolase; Protease.

SQ SEQUENCE 501 AA; 55816 MW; C085531345E024E CRC64;  
 Query Match 100.0%; Score 144; DB 2; Length 501;  
 Best Local Similarity 100.0%; Pred. No. 2,4e-13;  
 Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTLNLIVDTGSSNFAV 28  
 |||||  
 DB 74 GYVEMTVGSPPTLNLIVDTGSSNFAV 101

RESULT 8  
 Q9USL1 PRELIMINARY; PRT; 532 AA.  
 DT 01-MAR-2000 (TrEMBLrel. 13, Created)  
 DT 01-OCT-2001 (TrEMBLrel. 18, Last sequence update)  
 DT 01-MAR-2004 (TrEMBLrel. 26, Last annotation update)  
 DE KIAA1149 protein (Fragment).  
 GN Name=KIAA1149;  
 OS Homo sapiens (Human).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Primates; Catarrhini; Homnidae; Homo.  
 OC NCBI\_TaxID=9606;  
 OX [1]  
 RN SEQUENCE FROM N.A.  
 RP TISSUE=Brain; N.A.  
 RC MEDLINE=20039618; PubMed=10574461;  
 RA Hirotsawa M., Nagase T., Ishikawa K., Kikuno R., Nomura N., Ohara O.,  
 RT "Characterization of cDNA clones selected by the Genemark analysis  
 RT from size-fractionated cDNA libraries from human brain.";  
 RL DNA Res. 6:3329-336(1999).  
 CC -1- SIMILARITY: Belongs to peptidase family A1.  
 DR EMBL; AB032975; BAA8463.2; -.  
 DR HSSP; P56817; 1PKN.  
 DR GO; GO:0005768; C:Endosome; ISS.  
 DR GO; GO:0005794; C:Golgi apparatus; ISS.  
 DR GO; GO:0016021; C:Integral to membrane; ISS.  
 DR GO; GO:0004190; F:Aspartic-type endopeptidase activity; ISS.  
 DR GO; GO:0006035; F:beta-amyloid metabolism; ISS.  
 DR GO; GO:0006009; P:membrane protein ectodomain proteolysis; ISS.  
 DR InterPro: IPR001461; Peptidase A1.  
 DR InterPro: IPR009119; Pept\_A1\_BACE1.  
 DR InterPro: IPR009007; Pept\_Aspartic.  
 DR InterPro: IPR001969; Pept\_Asp\_AS.  
 DR PRINTS; PR01816; BACE1.  
 DR PRINTS; PR00792; PEPSIN.  
 DR PROSITE; PS00141; ASP\_PROTEASE; 1.  
 KM Aspartyl protease; Hydrolyase; Protease.  
 FT NON TER 1  
 SQ SEQUENCE 532 AA; 58720 MW; 98B135D0D5FBD2E8 CRC64;

Query Match 100.0%; Score 144; DB 2; Length 532;  
 Best Local Similarity 100.0%; Pred. No. 2,6e-13;  
 Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTLNLIVDTGSSNFAV 28  
 |||||  
 DB 105 GYVEMTVGSPPTLNLIVDTGSSNFAV 132

RESULT 9  
 Q8C7R1 PRELIMINARY; PRT; 501 AA.  
 AC 08C7R1;  
 DT 01-MAR-2003 (TrEMBLrel. 23, Created)  
 DT 01-MAR-2003 (TrEMBLrel. 23, Last sequence update)  
 DT 01-MAR-2004 (TrEMBLrel. 26, Last annotation update)  
 DE Mus musculus 12 days embryo spinal cord cDNA, RIKEN full-length  
 DE enriched library, clone:CS30008K17 product:beta-site App cleaving  
 DE enzyme, full insert sequence.

GN Name=Bacel; Synonyms=Bace;  
 OS Mus musculus (Mouse).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.  
 OC NCBI\_TaxID=10090;  
 RN [1]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Spinal cord;  
 RX MEDLINE=99279253; PubMed=10349636; DOI=10.1016/S0076-6879(99)03004-9;  
 RA Carninci P., Hayashizaki Y.;  
 RT "High-efficiency full-length cDNA cloning.";  
 RL Meth. Enzymol. 303:19-44(1999).  
 RN [2]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Spinal cord;  
 RX MEDLINE=21085660; PubMed=11217851; DOI=10.1038/35055500;  
 RA RIKEN FANTOM Consortium;  
 RT "Functional annotation of a full-length mouse cDNA collection.";  
 RL Nature 409:685-690(2001).  
 RN [3]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Spinal cord;  
 RA The FANTOM Consortium;  
 RT "Analysis of the mouse transcriptome based on functional annotation of  
 RT 60,770 full-length cDNAs.";  
 RL Nature 420:563-573(2002).  
 RN [4]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Spinal cord;  
 RX MEDLINE=20499374; PubMed=11042159; DOI=10.1101/9r.145100;  
 RA Carninci P., Shibata Y., Hayatsu N., Sugahara Y., Shibata K., Itoh M.,  
 RA Kono H., Okazaki Y., Muramatsu M., Hayashizaki Y.;  
 RT "Normalization and subtration of cap-trapper-selected cDNAs to  
 RT prepare full-length cDNA libraries for rapid discovery of new genes.";  
 RL Genome Res. 10:1617-1630(2000).  
 RN [5]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Spinal cord;  
 RX MEDLINE=20530913; PubMed=11076861; DOI=10.1101/gr.152600;  
 RA Shibata K., Itoh M., Aizawa K., Nagaoka S., Sasaki N., Carninci P.,  
 RA Kono H., Akiyama J., Nishi K., Katsunai T., Tashiro H., Itoh M.,  
 RA Sumi N., Ishii Y., Nakamura S., Hazama M., Nishine T., Harada A.,  
 RA Yamamoto R., Matsumoto H., Sakaguchi S., Ikegami T., Kashiwagi K.,  
 RA Fujiwaka S., Inoue K., Ogawa Y., Izawa M., Ohara E., Wataniki M.,  
 RA Yoneda Y., Ishikawa T., Ozawa K., Tanaka T., Matsura S., Kawai J.,  
 RA Okazaki Y., Muramatsu M., Inoue Y., Kira A., Hayashizaki Y.;  
 RT "RIKEN integrated sequence analysis (RISA) system-384-Format  
 RT sequencing pipeline with 384 multicapillary sequencer.";  
 RL Genome Res. 10:1757-1771(2000).  
 RN [6]  
 RP SEQUENCE FROM N.A.  
 RC STRAIN=C57BL/6J; TISSUE=Spinal cord;  
 RA Adachi J., Aizawa K., Akimura T., Arakawa T., Bono H., Carninci P.,  
 RA Fukuda S., Furuno M., Hanagaki T., Hara A., Hashizume W.,  
 RA Hayashida K., Hayatsu N., Hiramoto K., Hiraoka T., Hirozane T.,  
 RA Hori F., Imctani K., Ishii Y., Itoh M., Kogawa I., Kasukawa T.,  
 RA Katon H., Kawai J., Kojima Y., Kondo S., Kono H., Kouda M., Koya S.,  
 RA Kurihara C., Matsuyama T., Miyazaki A., Murata N., Nakamura M.,  
 RA Nishi K., Nomura K., Numazaki R., Ohno M., Ohnato N., Okazaki Y.,  
 RA Saito R., Saitoh H., Sakai C., Sakai K., Sakazume N., Sano H.,  
 RA Sasaki D., Shibata K., Shinagawa A., Shiraki T., Sogabe Y., Tagami M.,  
 RA Tagawa A., Takahashi F., Takaku-Akahira S., Takeda Y., Tanaka T.,  
 RA Tomaru A., Toyota T., Yasunishi A., Muramatsu M., Hayashizaki Y.;  
 RL Submitted (JUL-2001) to the EMBL/GenBank/DBJ databases.  
 CC -1- SIMILARITY: Belongs to peptidase family A1.  
 DR EMBL; AK049626; BAC33844.1; -.  
 DR HSSP; P56817; 1PKN.  
 DR MGD; MGI:1346542; Bacel.  
 DR GO; GO:0005768; C:Endosome; ISS.  
 DR GO; GO:0005615; C:extracellular space; TAS.  
 DR GO; GO:0005794; C:Golgi apparatus; ISS.  
 DR GO; GO:0016021; C:Integral to membrane; ISS.





Submitted (AUG-2003) to the EMBL/GenBank/DBJ databases.  
 RL EMBL: BC055989; AAH55989.1; -  
 DR HSSP; P56817; 1FKN.  
 DR GO; GO:0004194; F:pepsin A activity; IEA.  
 DR GO; GO:0008233; F:peptidase activity; IEA.  
 DR GO; GO:0006508; P:proteolysis and peptidolysis; IEA.  
 DR InterPro; IPR009007; Pept\_AspArtic.  
 DR InterPro; IPR001969; Pept\_Asp\_AS.  
 DR PROSITE; PS00141; ASP\_PROTEASE; 2.  
 DR SEQUENCE 500 AA; 54722 MW; 10F16756CAFDD0B CRC64;  
 Query Match 84.7%; Score 122; DB 2; Length 500;  
 Best Local Similarity 78.6%; Pred. No. 6e-10;  
 Matches 22; Conservative 4; Mismatches 2; Indels 0; Gaps 0;  
 Qy 1 GYVENTVGSPPQTNIIVDTGSSNFAY 28  
 Db 75 GYVELLIGSPQKNVILVDTGSSNFAY 102  
 RESULT 12  
 Q6PB20 PRELIMINARY; PRT; 499 AA.  
 AC Q6PB20;  
 DT 05-JUL-2004 (TrEMBLrel. 27, Created)  
 DT 05-JUL-2004 (TrEMBLrel. 27, Last sequence update)  
 DT 05-JUL-2004 (TrEMBLrel. 27, Last annotation update)  
 DE MGC68482 protein.  
 GN Name=MGC68482;  
 OS Xenopus laevis (African clawed frog).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Amphibia; Batrachia; Anura; Mesobatrachia; Pipidoidea; Pipidae;  
 OC Xenopodinae; Xenopus.  
 OX NCBI\_TaxID=83355;  
 OX [1]  
 RP SEQUENCE FROM N.A.  
 RC TISSUE=Kidney;  
 RX MEDLINE=22388257; PubMed=12477932; DOI=10.1073/pnas.242603899;  
 RA Strausberg R.L., Feingold E.A., Grouse L.H., Derge J.G.,  
 RA Klausner R.D., Collins F.S., Wagner L., Shennan C.M., Schuler G.D.,  
 RA Altschul S.F., Zeeberg B., Buetow K.H., Schaefer C.F., Bhat N.K.,  
 RA Hopkins R.F., Jordan H., Moore T., Max S.I., Wang J., Hsieh P.,  
 RA Diatchenko L., Marusina K., Farmer A.A., Rubin G.M., Hong L.,  
 RA Stapleton M., Soares M.B., Bonaldi W.F., Casavant T.L., Scheetz T.E.,  
 RA Brownstein M.J., Ustun T.B., Toshlyuk S., Carninci P., Prange C.,  
 RA Raha S.S., Loquellano N.A., Peters G.J., Abramson R.D., Mullan S.J.,  
 RA Bosak S.A., McGowan P.J., McKernan K.J., Malek J.A., Gunaratne P.H.,  
 RA Richards S., Morley K.C., Hale S., Garcia A.M., Gay L.J., Hulyk S.W.,  
 RA Villalón D.K., Muzny D.M., Sodergren E.J., Lu X., Gibbs R.A.,  
 RA Fahy J., Helton E., Kettelman M., Madan A., Rodriguez S., Sanchez A.,  
 RA Blakeley R.W., Touchman J.W., Green E.D., Dickson M.C.,  
 RA Rodriguez A.C., Grimwood J., Schmutz J., Myers R.M., Butterfield Y.S.,  
 RA Krzywinski M.I., Skalska U., Smalios D.E., Schnerch A., Schein J.E.,  
 RA Jones S.J., Marra M.A.,  
 RT "Generation and initial analysis of more than 15,000 full-length human  
 RT and mouse cDNA sequences."  
 RT Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903(2002).  
 RN [12]  
 RP SEQUENCE FROM N.A.  
 RC TISSUE=Kidney;  
 RX MEDLINE=22341132; PubMed=12454917; DOI=10.1002/dvdy.10174;  
 RA Klein S.L., Strausberg R.L., Wagner L., Pontius J., Clifton S.W.,  
 RA Richardson P.,  
 RT "Genetic and genomic tools for Xenopus research: The NIH Xenopus  
 RT initiative."  
 RT Dev. Dyn. 225:384-391(2002).  
 RN [13]  
 RP SEQUENCE FROM N.A.  
 RC TISSUE=Kidney;  
 RX Klein S., Strausberg R.,  
 RT Submitted (OCT-2003) to the EMBL/GenBank/DBJ databases.  
 CC -1- SIMILARITY: Belongs to peptidase family A1.

DR EMBL: BC059963; AAH59963.1; -  
 DR HSSP; P20142; 1AVF.  
 DR GO; GO:0009049; F:aspartic-type signal peptidase activity; IEA.  
 DR GO; GO:0004194; F:pepsin A activity; IEA.  
 DR GO; GO:0008233; F:peptidase activity; IEA.  
 DR GO; GO:0006508; P:proteolysis and peptidolysis; IEA.  
 DR InterPro; IPR001461; Peptidase\_A1.  
 DR InterPro; IPR009119; Pept\_A1\_BACE.  
 DR InterPro; IPR009121; Pept\_A1\_BACE2.  
 DR InterPro; IPR009007; Pept\_AspArtic.  
 DR InterPro; IPR001969; Pept\_Asp\_AS.  
 DR PRINTS; PR01817; BACE2.  
 DR PRINTS; PR01815; BACEFAMILY.  
 DR PRINTS; PR00792; PEPsin.  
 DR PROSITE; PS00141; ASP\_PROTEASE; 2.  
 DR Aspartyl protease; Hydrolyase; Protease.  
 DR SEQUENCE 499 AA; 54803 MW; E846674A5DF68AF CRC64;  
 Query Match 83.3%; Score 120; DB 2; Length 499;  
 Best Local Similarity 75.0%; Pred. No. 1.2e-09;  
 Matches 21; Conservative 5; Mismatches 2; Indels 0; Gaps 0;  
 Qy 1 GYVENTVGSPPQTNIIVDTGSSNFAY 28  
 Db 74 GYVELLIGTTPQKNVILVDTGSSNFAY 101  
 RESULT 13  
 Q9NZL1 PRELIMINARY; PRT; 396 AA.  
 AC Q9NZL1;  
 DT 01-OCT-2000 (TrEMBLrel. 15, Created)  
 DT 01-OCT-2000 (TrEMBLrel. 15, Last sequence update)  
 DT 01-MAR-2004 (TrEMBLrel. 26, Last annotation update)  
 DE Aspartyl protease.  
 GN Name=BACE2;  
 OS Homo sapiens (Human).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.  
 OX NCBI\_TaxID=9606;  
 OX [1]  
 RP SEQUENCE FROM N.A.  
 RX MEDLINE=20422477; PubMed=10965118;  
 RA Solans A., Estivill X., de la Luna S.,  
 RT "A new aspartyl protease on 21q22.3, BACE2, is highly similar to  
 RT Alzheimer's amyloid precursor protein beta-secretase."  
 RT Cytogenet. Cell Genet. 89:177-184(2000).  
 CC -1- SIMILARITY: Belongs to peptidase family A1.  
 DR EMBL; AF188277; AAF35836.1; -  
 DR HSSP; P56817; 1FKN.  
 DR GO; GO:0016021; C:integral to membrane; ISS.  
 DR GO; GO:0006509; P:membrane protein ectodomain proteolysis; ISS.  
 DR GO; GO:0043985; P:negative regulation of amyloid precursor pr. . .; ISS.  
 DR GO; GO:0016486; P:peptide hormone processing; ISS.  
 DR InterPro; IPR001461; Peptidase\_A1.  
 DR InterPro; IPR009119; Pept\_A1\_BACE.  
 DR InterPro; IPR009121; Pept\_A1\_BACE2.  
 DR InterPro; IPR009007; Pept\_AspArtic.  
 DR InterPro; IPR001969; Pept\_Asp\_AS.  
 DR PRINTS; PR01815; BACE2.  
 DR PRINTS; PR00792; PEPsin.  
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 DR Aspartyl protease; Hydrolyase; Protease.  
 DR SEQUENCE 396 AA; 43013 MW; 5023A7AF391CEAC9 CRC64;  
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 Best Local Similarity 78.6%; Pred. No. 1.3e-09;  
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 Qy 1 GYVENTVGSPPQTNIIVDTGSSNFAY 28  
 Db 91 GYVELLIGTTPQKNVILVDTGSSNFAY 118







CC therapy. (1) can be used for producing preparations of homogeneity  
CC processed BACE that may be used for e.g. studying or treating diseases  
CC such as Alzheimer's disease or Down's syndrome. The human BACE1 gene is  
CC located on chromosome 11, more specifically to 11q23.2-23.3. The present  
CC sequence represents a human BACE1 isoform A protease domain amino acid  
CC sequence, which is used in the exemplification of the present invention.  
XX  
SQ Sequence 28 AA;  
  
Query Match 100.0%; Score 144; DB 8; Length 28;  
Best Local Similarity 100.0%; Pred. No. 6,9e-15;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
  
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Dd 1 GYVYEMTVGSPPTLTNLIVDGSNFAV 28  
  
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ID AAU23068 standard; protein; 387 AA.  
XX  
AC AAU23068;  
XX  
DT 17-DEC-2001 (first entry)  
XX  
DE Novel human enzyme polypeptide #154.  
XX  
KX Human; oxidoreductase enzyme; transferase; hydrolase; lyase; isomerase;  
KX ligase; hyperproliferative disorder; immunodeficiency disorder;  
KM autoimmune disorder; neurological disorder; metabolic disorder;  
KM inflammatory disorder; cardiovascular disorder; reproductive disorder;  
KM blood-related disorder; infectious disorder; cytostatic; anti arthritic;  
KM nephrotropic; anticoagulant.  
XX  
OS Homo sapiens.  
XX  
PN WO200155301-A2.  
XX  
PD 02-AUG-2001.  
XX  
PF 17-JAN-2001; 2001WO-US001239.  
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PR 05-JAN-2001; 2001US-0259678P.  
XX  
XX (HUMA-) HUMAN GENOME SCI INC.  
XX  
PI Rosen CA, Barash SC, Ruben SM;  
XX  
XX WPI; 2001-465566/50.  
XX N-PSDB; AAS40938.  
XX  
PT Novel polypeptides and polynucleotides useful for diagnosing, preventing,  
XX treating neural, immune system, muscular, reproductive, pulmonary,  
XX cardiovascular, renal, proliferative disorders and cancerous diseases.  
XX  
PS Claim 11; SEQ ID NO 1064; 1180bp; English.  
XX  
XX The present invention relates to the isolation of novel human enzyme  
XX polypeptides, and the cDNA (AAS40795-AAS41684) and genomic sequences  
XX encoding them. The enzyme polypeptides of the invention may comprise the  
XX functional classes of oxidoreductases, transferases, hydrolases, lyases,  
XX isomerases or ligases. The sequences of the invention are useful in the  
XX diagnosis, treatment, prevention and/or prognosis of a wide range of  
XX disorders including hyperproliferative disorders (e.g. cancer),  
XX immunodeficiency disorders (e.g. AIDS) autoimmune disorders (e.g.  
XX arthritis), neurological disorders (e.g. Alzheimer's disease), metabolic  
XX disorders (e.g. phenylketonuria), inflammatory disorders (e.g. asthma),  
XX cardiovascular disorders (e.g. atherosclerosis), blood-related disorders  
XX (e.g. haemophilia), reproductive disorders (e.g. infertility) and  
XX infectious disorders (e.g. influenza). The polynucleotides of the  
XX invention can also be used in gene therapy. AAU22915-AAU23814 represent  
XX the novel human enzyme polypeptides of the invention. Note: The sequence  
XX data for this patent did not form part of the printed specification, but  
XX was obtained in electronic format directly from WIPO at  
XX ftp.wipo.int/pub/published\_pct\_sequences  
XX  
SQ Sequence 387 AA:  
XX  
XX Query Match 100.0%; Score 144; DB 4; Length 387;  
XX Best Local Similarity 100.0%; Pred. No. 1,6e-13;  
XX Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
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RESULT 3  
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XX  
XX AAU23069;  
AC  
XX 17-DEC-2001 (first entry)  
DT  
XX  
XX Novel human enzyme polypeptide #155.  
DE  
XX Human; oxidoreductase enzyme; transferase; hydrolase; lyase; isomerase;  
XX ligase; hyperproliferative disorder; immunodeficiency disorder;  
XX autoimmune disorder; neurological disorder; metabolic disorder;  
XX inflammatory disorder; cardiovascular disorder; reproductive disorder;  
XX blood-related disorder; infectious disorder; cytostatic; anti arthritic;  
XX nephrotropic; anticoagulant.  
XX  
XX Homo sapiens.  
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XX 17-JUN-2001; 2001WO-US001239.  
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PR 11-DEC-2000; 2000US-0254097P.  
PR 05-JAN-2001; 2001US-0259678P.  
PA (HUMAN) HUMAN GENOME SCI INC.  
PI Rosen CA, Barash SC, Ruben SM;  
XX WPI; 2001-465566/50.  
DR N-PSDB; AAS40939.  
XX  
PT Novel polypeptides and polynucleotides useful for diagnosing, preventing,  
PT treating neutral, immune system, muscular, reproductive, pulmonary,  
PT cardiovascular, renal, proliferative disorders and cancerous diseases.  
PS  
XX Claim 11; SEQ ID NO 1065; 1180bp; English.  
XX  
CC The present invention relates to the isolation of novel human enzyme  
CC polypeptides, and the cDNA (AAS40785-AAS41684) and genomic sequences  
CC encoding them. The enzyme polypeptides of the invention may comprise the  
CC functional classes of oxidoreductases, transferases, hydrolases, lyases,  
CC isomerases or ligases. The sequences of the invention are useful in the  
CC diagnosis, treatment, prevention and/or prognosis of a wide range of  
CC disorders including hyperproliferative disorders (e.g. cancer),  
CC immunodeficiency disorders (e.g. AIDS), autoimmune disorders (e.g.  
CC arthritis), neurological disorders (e.g. Alzheimer's disease), metabolic  
CC disorders (e.g. phenylketonuria), inflammatory disorders (e.g. asthma),  
CC cardiovascular disorders (e.g. atherosclerosis), blood-related disorders  
CC (e.g. haemophilia), reproductive disorders (e.g. infertility) and  
CC infectious disorders (e.g. influenza). The polynucleotides of the  
CC invention can also be used in gene therapy. AAU22915-AAU23814 represent  
CC the novel human enzyme polypeptides of the invention. Note: The sequence  
CC data for this patent did not form part of the printed specification, but  
CC was obtained in electronic format directly from WIPO at  
CC ftp.wipo.int/pub/published\_pct\_sequences  
XX  
SQ Sequence 390 AA;  
SQ  
Query Match 100.0%; Score 144; DB 4; Length 390;  
Best Local Similarity 100.0%; Pred. No. 1,6e-13;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
CY 1 GYVEMTVGSPPTLNTILVDTGSSNFV 28  
Db 13 GYVEMTVGSPPTLNTILVDTGSSNFV 40  
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RESULT 4  
ID ADC81581 standard; protein; 391 AA.  
XX  
AC ADC81581;  
XX  
DT 01-JAN-2004 (first entry)  
XX  
DE Beta-secretase (1fkn) amino acid sequence SEQ ID NO:4.  
XX

KM	human; BACE; modification; Pro33lys; pro-enzyme.
XX	Unidentified.
OS	
XX	
PN	MO2003072733-A2.
PD	04-SEP-2003.
XX	
XX	21-FEB-2003; 2003WO-US005508.
PP	
XX	21-FEB-2002; 2002US-0358651P.
PR	
XX	(PHAA ) PHARMACIA & UPJOHN CO.
PA	
XX	
P1	Chou K, Howe JW;
DR	WPI; 2003-712719/67.
XX	
PT	BACE polypeptides having Pro33lys modification, useful in determining possible mutations, which will inhibit enzyme activity, and in determining potential active site for target molecules.
XX	
PS	Disclosure; Fig 3; 38pp; English.
XX	
CC	The present invention describes an isolated polypeptide (I) comprising or consisting of a fully defined sequence of 432 amino acids (see ADG81561), CC and comprising human BACE having the modification Pro33lys. Also described: (1) a composition comprising an active human BACE enzyme CC comprising the pro-enzyme sequence of BACE having the modification Pro33lys; (2) an isolated polynucleotide comprising a sequence encoding CC (1); (3) an isolated polynucleotide consisting of or comprising of CC nucleotides 70-1365 of a 1355-bp sequence (see ADG81562); (4) an CC expression vector comprising the polynucleotide of (2), or a CC polynucleotide sequence encoding a Pro33lys-BACE polypeptide, where the CC expression vector can produce the Pro33lys-BACE polypeptide when present in a compatible host cell, when cultured under conditions that allow CC production; (5) a recombinant host cell comprising the expression vector; CC and (6) producing a (active) Pro33lys-BACE polypeptide. The BACE CC polypeptide having Pro33lys modification may be used in determining CC possible mutations, which will inhibit enzyme activity, and in CC determining potential active site for target molecules. The vector CC comprising the BACE polynucleotide is useful for producing recombinant CC BACE polypeptides having Pro33lys modification. The present sequence CC represents a beta-secretase amino acid sequence, which is used in the CC exemplification of the present invention.
SQ	Sequence 391 AA;
Query Match	100.0%; Score 144; DB 7; Length 391;
Best Local Similarity	100.0%; Pred. No. 1.6e-13;
Matches	28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
OY	1 GYYEMTVGSPQTINILVDTGSSNFAV 28       IDB 19 GYYEMTVGSPQTINILVDTGSSNFAV 46
Db	
RESULT 5	
KM	AD164643 standard; protein; 403 AA.
ID	AD164643
XX	
XX	AD164643;
AC	
XX	
DT	22-APR-2004 (first entry)
XX	
DE	Mature human beta-secretase (BACE) protein seq id 4.
XX	
KM	cysatal; glycosylated human beta-secretase; BACE; human beta-secretase; protein co-ordinate data.
XX	
OS	Homo sapiens.
XX	
PN	US2004014194-A1.

```

XX 22-JAN-2004.
XX
XX 26-MAR-2003; 2003US-00400273.
XX
XX 27-MAR-2002; 2002US-0367937P.
XX
XX (SCHE ) SCHERING CORP.
XX
XX Beyer BM, Hammond GS, Reichert P, Strickland C, Wang W, Weber PC;
XX Wong GT, Zhang L;
XX
XX WPI: 2004-167920/16.
XX
XX
XX Claim 5; SEQ ID NO 4; 107pp; English.
XX
XX The invention describes a crystal comprising a glycosylated, human beta-
XX secretase polypeptide characterised by structural coordinates comprising
XX a root mean square deviation of conserved residue backbone atoms of less
XX than 1.5 Angstrom when superimposed on backbone atoms described by
XX structural coordinates. The crystal is useful for determining the three-
XX dimensional structure of beta-secretase and other related proteins. This
XX is the amino acid sequence of a mature human beta-secretase (BACE)
XX protein.
XX
XX Sequence 403 AA;
XX
XX Query Match 100.0%; Score 144; DB 8; Length 403;
XX Best Local Similarity 100.0%; Pred. No. 1.7e-13;
XX Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
XX
XX 1 GYVEMTWGSPPTLNLIVDGSNPAV 28
XX |||||
XX 27 GYVEMTWGSPPTLNLIVDGSNPAV 54
XX
XX RESULT 6
XX ADI64644
XX ID ADI64644 standard; protein; 408 AA.
XX
XX AC ADI64644;
XX
XX DT 22-APR-2004 (first entry)
XX
XX DE Mature human beta-secretase (BACE) protein seq id 5.
XX
XX KW crystal; glycosylated human beta-secretase; BACE; human beta-secretase;
XX protein co-ordinate data.
XX
XX OS Homo sapiens.
XX
XX PN US2004014194-A1.
XX
XX PD 22-JAN-2004.
XX
XX PF 26-MAR-2003; 2003US-00400273.
XX
XX PR 27-MAR-2002; 2002US-0367937P.
XX
XX PA (SCHE ) SCHERING CORP.
XX
XX PI Beyer BM, Hammond GS, Reichert P, Strickland C, Wang W, Weber PC;
XX Wong GT, Zhang L;
XX
XX WPI: 2004-167920/16.
XX
XX New crystal comprising a glycosylated, human beta-secretase polypeptide,
XX useful for determining the three-dimensional structure of beta-secretase
XX and other related proteins.

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OS Synthetic.  
 XX  
 PN WO2004011641-A2.  
 XX  
 PD 05-FEB-2004.  
 XX  
 PP 25-JUL-2003; 2003WO-GB003200.  
 XX  
 PR 26-JUL-2002; 2002US-0398681P.  
 XX  
 PA (ASTE-) ASTEX TECHNOLOGY LTD.  
 XX  
 PI Vuillard LM, Patel SJ, Yon JR, Cleasby A, Hamilton BJ, Shah A;  
 XX  
 DR WPI, 2004-169242/16.  
 XX  
 PT New beta site APP cleaving enzyme (BACE) protein, useful for treating or  
 PT preventing Alzheimer's disease or Alzheimer's-type pathology of Down's  
 PT syndrome.  
 XX  
 PS Claim 10; SEQ ID NO 20; 145pp; English.  
 XX  
 CC The present invention relates to a beta site APP cleaving enzyme (BACE)  
 CC protein. The compound or the composition is useful in medicine and the  
 CC BACE crystal structure is useful for drug discovery. The BACE protein,  
 CC compounds, pharmaceutical compositions, medicament, drug or other  
 CC composition comprising the compound is useful for treating or preventing  
 CC Alzheimer's disease or Alzheimer's-type pathology of Down's syndrome. The  
 CC present sequence represents the DNA sequence for a BACE protein.  
 CC  
 SQ Sequence 411 AA;

Query Match 100.0%; Score 144; DB 8; Length 411;  
 Best Local Similarity 100.0%; Pred. No. 1.7e-13;  
 Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVENTVGSPPQTILNIVDTGSSNPAV 28  
 DB 32 GYVENTVGSPPQTILNIVDTGSSNPAV 59

RESULT 10  
 ABR61930  
 ID ABR61930 standard; protein; 414 AA.  
 XX  
 AC ABR61930;  
 XX  
 DT 12-SEP-2003 (first entry)  
 XX  
 DE Human promemapsin 2 protein fragment.  
 XX  
 KW Memapsin 1; nootropic; neuroprotective; memapsin 2; beta secretase;  
 KW beta-amyloid protein; Alzheimer's disease; promemapsin 2; human.  
 XX  
 OS Homo sapiens.  
 XX  
 PN WO2003039454-A2.  
 XX  
 PD 15-MAY-2003.  
 XX  
 PP 23-OCT-2002; 2002WO-US043324.  
 XX  
 PR 23-OCT-2001; 2001US-0335952P.  
 PR 27-NOV-2001; 2001US-0333545P.  
 PR 14-JAN-2002; 2002US-0348464P.  
 PR 14-JAN-2002; 2002US-0348615P.  
 PR 20-JUN-2002; 2002US-0390804P.  
 PR 19-JUL-2002; 2002US-0397557P.  
 PR 19-JUL-2002; 2002US-0397619P.  
 XX  
 PA (OKLA-) OKLAHOMA MEDICAL RES FOUND.  
 PA (UNIT ) UNIV ILLINOIS FOUND.  
 XX

PI Ghosh AK, Tang J, Bilcer G, Chang W, Hong L, Koelsch G, Loy J;  
 PI Turner RT;  
 XX  
 DR WPI; 2003-541410/51.  
 XX  
 PP New peptide compounds are memapsin beta secretase inhibitors used for  
 PT treating Alzheimer's disease.  
 XX  
 PS Example; Fig 12; 407pp; English.  
 XX  
 CC The invention relates to peptide compounds of specified formula. The  
 CC compounds exhibit memapsin 2-beta secretase inhibitory activity relative  
 CC to memapsin 1-beta secretase and reduce the accumulation of beta-amyloid  
 CC protein. The compounds can be used for treating Alzheimer's disease. The  
 CC present sequence represents a human promemapsin 2 protein fragment used  
 CC in crystal structures  
 XX  
 SQ Sequence 414 AA;

Query Match 100.0%; Score 144; DB 6; Length 414;  
 Best Local Similarity 100.0%; Pred. No. 1.7e-13;  
 Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVENTVGSPPQTILNIVDTGSSNPAV 28  
 DB 34 GYVENTVGSPPQTILNIVDTGSSNPAV 61

RESULT 11  
 ADC72735  
 ID ADC72735 standard; protein; 414 AA.  
 XX  
 AC ADC72735;  
 XX  
 DT 18-DEC-2003 (first entry)  
 XX  
 DE Human beta-site aspartyl protease cleaving enzyme catalytic domain.  
 XX  
 KW neuroprotective; nootropic; crystalline; Beta-site APP cleaving enzyme;  
 KW BACE; aspartyl protease; Alzheimer's disease; protein co-ordinate data.  
 XX  
 OS Homo sapiens.  
 XX  
 PN WO2003012089-A2.  
 XX  
 PD 13-FEB-2003.  
 XX  
 PF 26-JUL-2002; 2002WO-GB003461.  
 XX  
 PR 26-JUL-2001; 2001US-0308366P.  
 XX  
 PA (ASTE-) ASTEX TECHNOLOGY LTD.  
 PA (JANC ) JANSSEN PHARM NV.  
 XX  
 PI Yon J, Cleasby A, Bruinzeel WD, Masure SLJ, Tickle I, Sharff A;  
 XX  
 DR WPI; 2003-239524/23.  
 DR N-PSDB; ADC72736.  
 XX  
 PT New Beta-site APP cleaving enzyme (BACE) proteins and protein crystal,  
 PT useful in designing compounds that inhibit or modulate BACE, in drug  
 PT screening assays, and in identifying receptors.  
 XX  
 PS Disclosure; Fig 2a; 272pp; English.  
 XX  
 CC The invention relates to a new crystalline form of Beta-site APP cleaving  
 CC enzyme (BACE) or its functional portion having an active site containing  
 CC one or more ligands other than the natural substrate or the substrate  
 CC that occurs naturally or physiologically within the active site.  
 CC Inhibitors of BACE protein or its functional portion is useful for  
 CC preparing a composition or medicament for inhibiting BACE or the  
 CC production of A-beta or its fragments, and in therapy for treating  
 CC Alzheimer's disease. The BACE crystals and proteins may be used to design

CC compounds that inhibit or modulate BACE, in drug screening assays, and in  
CC identifying receptors. This sequence represents a fragment of the full  
CC length BACE protein from amino acid 76 to the C-terminus.  
XX  
SQ Sequence 414 AA;

Query Match 100.0%; Score 144; DB 7; Length 414;  
Best Local Similarity 100.0%; Pred. No. 1.7e-13;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTQNLIVDTGSSNFAV 28  
DB 29 GYVEMTVGSPPTQNLIVDTGSSNFAV 56

RESULT 12  
AAB07899  
ID AAB07899 standard; protein; 415 AA.  
XX  
AC AAB07899;  
XX  
DT 14-NOV-2000 (first entry)  
XX

DE Amino acid sequence of a human beta-secretase enzyme fragment.  
XX  
KW Beta-secretase; beta-amyloid precursor protein; beta-amyloid peptide;  
KW amyloid plaque component; Alzheimer's disease; amyloidogenic disease;  
KW inhibitor.  
XX  
OS Homo sapiens.  
XX  
FN WO200047618-A2.  
XX  
PD 17-AUG-2000.  
XX  
PF 10-FEB-2000; 2000WO-US003819.  
XX  
PR 10-FEB-1999; 99US-0119571P.  
PR 15-JUN-1999; 99US-0139172P.  
XX  
PA (ELAN-) ELAN PHARM INC.  
XX  
PI Anderson JP, Basl G, Doane MT, Frigon N, John V, Power M;  
PI Sima S, Tatsuno G, Tung J, Wang S, McConlogue L;  
XX  
DR WPI; 2000-533011/48.  
XX

PT Purified beta-secretase protein used in assays to discover inhibitors  
PT which can be used for the treatment of amyloidogenic diseases e.g.  
PT Alzheimer's disease.  
XX  
XX Claim 10; Fig 3B; 121pp; English.  
XX

PS The specification describes a beta-secretase enzyme. The enzyme cleaves  
CC beta-amyloid precursor protein to produce beta-amyloid peptide. This  
CC enzyme is therefore implicated in the production of amyloid plaque  
CC components which accumulate in the brains of individuals afflicted with  
CC Alzheimer's disease. Inhibitors of beta-secretase are administered to a  
CC mammalian subject e.g. with Alzheimer's disease or Alzheimer's disease-  
CC like pathology to test if they maintain or improve cognitive ability or  
CC reduce the plaque burden. The compounds are used for the treatment of  
CC amyloidogenic diseases e.g. Alzheimer's disease. The present sequence  
CC represents a human beta-secretase enzyme fragment  
XX  
SQ Sequence 415 AA;

Query Match 100.0%; Score 144; DB 3; Length 415;  
Best Local Similarity 100.0%; Pred. No. 1.8e-13;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTQNLIVDTGSSNFAV 28  
DB 29 GYVEMTVGSPPTQNLIVDTGSSNFAV 56

RESULT 13  
ADJ57792  
ID ADJ57792 standard; protein; 417 AA.  
XX  
AC ADJ57792;  
XX  
DT 06-MAY-2004 (first entry)  
XX

DE BACE N-Q R56RR57 crystallised protein.  
XX  
KW beta site APP cleaving enzyme; BACE; Nootropic; Neuroprotective;  
KW Alzheimer's disease.  
XX  
OS Synthetic.  
XX  
FN WO2004011641-A2.  
XX  
PD 05-FEB-2004.  
XX  
PF 25-JUL-2003; 2003WO-GB003200.  
XX  
PR 26-JUL-2002; 2002US-0398681P.  
XX  
PA (ASTE-) ASTEX TECHNOLOGY LTD.  
XX  
PI Vulliamd LMM, Patel SJ, Yon JR, Cleasby A, Hamilton BJ, Shah A;  
XX  
DR WPI; 2004-169242/16.  
XX

PT New beta site APP cleaving enzyme (BACE) protein, useful for treating or  
PT preventing Alzheimer's disease or Alzheimer's-type pathology of Down's  
PT syndrome.  
XX  
PS Claim 10; SEQ ID NO 21; 145pp; English.  
XX  
XX The present invention relates to a beta site APP cleaving enzyme (BACE)  
CC protein. The compound or the composition is useful in medicine and the  
CC BACE crystal structure is useful for drug discovery. The BACE protein,  
CC compounds, pharmaceutical compositions, medication, drug or other  
CC composition comprising the compound is useful for treating or preventing  
CC Alzheimer's disease or Alzheimer's-type pathology of Down's syndrome. The  
CC present sequence represents the DNA sequence for a BACE protein.  
XX  
SQ Sequence 417 AA;

Query Match 100.0%; Score 144; DB 8; Length 417;  
Best Local Similarity 100.0%; Pred. No. 1.8e-13;  
Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 GYVEMTVGSPPTQNLIVDTGSSNFAV 28  
DB 32 GYVEMTVGSPPTQNLIVDTGSSNFAV 59

RESULT 14  
AAV88437  
ID AAV88437 standard; protein; 425 AA.  
XX  
AC AAV88437;  
XX  
DT 03-AUG-2000 (first entry)  
XX

DE Human Asp2 amino acid sequence containing proteolytic cleavage site.  
XX  
XX Aspartyl protease; aspartase; amyloid precursor protein; APP; Asp 2;  
KW Alzheimer's disease; beta secretase site.  
XX  
OS Homo sapiens.  
XX  
FN WO200017369-A2.  
XX



PD 30-MAR-2000.  
 XX  
 XX 23-SEP-1999; 99WO-US020861.  
 XX  
 XX 24-SEP-1998; 98US-0101594P.  
 XX  
 PA (PHMA ) PHARMACIA & UPJOHN CO.  
 XX  
 PI Gurney ME, Bienkowski MJ, Heinrichson RL, Parodi LA, Yan R;  
 XX  
 DR MPI; 2000-303209/26.  
 XX  
 DR N-PSDB; AAA15677.  
 XX  
 PT New enzyme designated human aspartase useful in research into Alzheimer's  
 PT Disease is capable of cleaving amyloid protein precursor at the beta  
 PT secretase site to produce amyloid beta peptide.  
 XX  
 PS Example 9; Page 166-168; 183pp; English.  
 XX  
 CC This sequence represents a modified version of the human aspartase 2  
 CC (Asp2) nucleotide sequence. The sequence is used in the bacterial  
 CC expression of human Asp2L. The invention relates to a protease (e.g.  
 CC Asp2) capable of cleaving the beta secretase site of amyloid precursor  
 CC protein (APP). The protease contains a sequence encoding the amino acid  
 CC sequence DTG and a sequence encoding DSG or DTG separated by 100-300  
 CC amino acids. When mutated the APP gene causes an autosomal dominant form  
 CC of Alzheimer's disease. APP localises to the cell surface membrane and  
 CC have a single C-terminal transmembrane domain. Proteolytic processing of  
 CC APP produces the amyloid beta protein, which is possibly very important  
 CC in Alzheimer's disease. The invention includes a nucleotide sequence  
 CC encoding the protease, a vector containing the nucleotide sequence, and a  
 CC cell line comprising the vector. Methods for screening for inhibitors of  
 CC beta secretase activity are also given in the invention. The human  
 CC aspartase protein and nucleotide sequences and the methods for  
 CC identifying inhibitors of the protease, are useful in the treatment of  
 CC and research in to Alzheimer's disease  
 XX  
 SO Sequence 425 AA;  
 XX  
 QY Query Match 100.0%; Score 144; DB 3; Length 425;  
 DB Best Local Similarity 100.0%; Pred. No. 1.8e-13;  
 Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 QY 1 GYVEMTVGSPPTNTLVDTGSSNFAV 28  
 DB 46 GYVEMTVGSPPTNTLVDTGSSNFAV 73  
 XX  
 RESULT 15  
 AAU07214  
 ID AAU07214 standard; protein; 425 AA.  
 XX  
 AC AAU07214;  
 XX  
 DT 11-SEP-2003 (revised)  
 DT 24-OCT-2001 (first entry)  
 XX  
 DE T7-caspase-caspase 8-human aspartyl protease 2a deltaTM.  
 XX  
 KM Human; aspartyl protease 1; Asp-1; neurotrophic; neuroprotective;  
 KM aspartyl protease 2; Asp2; amyloid protein precursor; APP;  
 KM beta-secretase; Alzheimer's disease.  
 XX  
 XX Homo sapiens.  
 OS Enterobacteria phage T7.  
 OS  
 PN MO200149097-A2.  
 XX  
 PD 12-JUL-2001.  
 PD  
 PF 09-MAY-2001; 2001MO-IB000797.  
 PF  
 XX 09-MAY-2001; 2001MO-IB000797.  
 XX

XX  
 XX (BIEN/) BIENKOWSKI M J.  
 PA (GURN/) GURNEY M E.  
 PA (HEIN/) HEINRICKSON R L.  
 PA (PARO/) PARODI L A.  
 PA (YANR/) YAN R.  
 XX  
 PI Bienkowski MJ, Gurney ME, Heinrichson RL, Parodi LA, Yan R;  
 XX  
 DR MPI; 2001-502548/55.  
 XX  
 DR N-PSDB; AAS11714.  
 XX  
 PT Novel purified polypeptide comprising fragment of mammalian aspartyl  
 PT protease 2, lacking Asp2 transmembrane domain and retaining beta  
 PT secretase activity of Asp2 useful for identifying inhibitors of Asp2  
 PT activity.  
 XX  
 PS Example 9; Page 158-159; 185pp; English.  
 XX  
 CC The invention relates to a novel purified polypeptide comprising a  
 CC fragment of mammalian aspartyl protease 2 (Asp2) protein which lacks the  
 CC Asp2 transmembrane domain and the Asp2 protein, and where the polypeptide  
 CC and the fragment retain the beta-secretase activity of the mammalian Asp2  
 CC protein. Also included is an isoform of amyloid protein precursor (APP)  
 CC comprising the amino acid sequence of a APP or its fragment containing an  
 CC APP cleavage site recognisable by a mammalian beta-secretase, and further  
 CC comprising two lysine residues at the carboxyl terminus of the amino acid  
 CC sequence of the mammalian APP or APP fragment. The polypeptides are used  
 CC for assaying for modulators of beta-secretase activity; identifying  
 CC agents that inhibit the APP processing activity of human Asp2 aspartyl  
 CC protease (Hu-Asp2); identifying agents that modulate the activity of Asp2  
 CC ; and for reducing cellular production of amyloid beta (Abeta) from APP.  
 CC Agents identified by the above methods are useful for treating  
 CC Alzheimer's disease; and for identifying modulators of amyloid-beta  
 CC (Abeta) peptide production, for use in designing therapeutics for the  
 CC treatment or prevention of Alzheimer's disease. Probes and primers  
 CC derived from Asp nucleic acid sequences are useful for detecting Hu-Asp  
 CC nucleic acids in in vitro assays and in Northern and Southern blots. The  
 CC present sequence represents the amino acid sequence of T7-caspase-caspase  
 CC 8-human-Asp-2a delta TM construct which has a T7 tag, a caspase 8 leader  
 CC sequence and cleavage site, and lacks the transmembrane domain. This  
 CC construct was used for bacterial expression and purification of human  
 CC Asp2a. (Updated on 11-SEP-2003 to standardise OS field)  
 XX  
 SO Sequence 425 AA;  
 XX  
 QY Query Match 100.0%; Score 144; DB 4; Length 425;  
 DB Best Local Similarity 100.0%; Pred. No. 1.8e-13;  
 Matches 28; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 QY 1 GYVEMTVGSPPTNTLVDTGSSNFAV 28  
 DB 46 GYVEMTVGSPPTNTLVDTGSSNFAV 73  
 XX

Search completed: July 26, 2005, 16:44:20  
 Job time : 166 secs

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